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gaccettgga atgccaagtt caagtttage tatgtetege ggagaggeeg gtggaagaag
                                                                      180
caacgagaat gaagcacccc agttctctgc tgagcacatg ggcatctgca ataaagattt
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aatttcccag cttctcctga agctcggtat ggccacaaca ctaaattctg cccgaggaga
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ttgagcaaaa tagtatggga cttccaagaa atg ttt tta aag tca ggg gca ggc
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                                 Met Phe Leu Lys Ser Gly Ala Gly
ctt tct tca tgc ctt ctt cct ctt tgc tgg ctg gaa cgc aaa gac cat
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Leu Ser Ser Cys Leu Leu Pro Leu Cys Trp Leu Glu Arg Lys Asp His
                        -5
ggc agg agg cca agc asc cat cct gga agg tgaaagcete atactaagga
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Gly Arg Arg Pro Ser Xaa His Pro Gly Arg
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cgtcaracag cgaaataara rcctgggtcc ttgaccctgt aaasatctcc ctccccatcc
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aaaaaaaaa aaa
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Leu	Ser	Arg		Glu	Cys	Val	His		Asp	Gly	Arg	Val		Thr	Leu	
			-35					-30					-25			202
tct	tat	cag	gag	cag	gag	cta	cag	gat	ttt	ctt	ctg	TCT	cag	atg	Cca	202
Ser	Tyr		Glu	Gln	Glu	Leu		Asp	Phe	Leu	Leu		GIN	met	ser	
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GIN		GIN	val	His	ATA	_	GIII	GIII	пеп	Ala	пур	vai	MEL	GIY	10	
	-5	a+a	200	ttc	3.C.t	1	cat	ata	aaa	÷++	~~=	cct	ata	aaa		298
Cla	yra Val	Lau	Cor	Phe	Cor	Acn	Wie	Val	61 v	Len	Glv	Dro	Tle	Glu	Ser	270
GIII	VQI	Deu	261	15	261	A311	1113	vui	20	neu	Cly	110	**	25	JUL .	
aht	aat	aat	aca	tct	acc	atc	acq	ata		ccc	caa	ata	ata		ato	346
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1144	017		30		••••			35					40			
cta	ttt	caq		gta	atq	qac	ctq		qtq	qca	qca	aga	tta	tgg	ttc	394
Leu	Phe	Gln	Phe	Val	Met	Asp	Leu	Lys	Val	Ala	Åla	Arg	Leu	Trp	Phe	
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Ser	Phe	Leu	Val	Thr	Asn	Val	Lys	Thr	Phe	Gln	Lys	Val	Met	Phe	Tyr	
	60					65					.70					
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Lys	Ile	Thr	Asn	Gly	Val	Ile	Phe	Val	Gly		Ser	Lys	Lys	Phe		
75					80					85					90	
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Gly	Ile	Lys	Trp	Lys	Val	Xaa	Ile	Leu		He	Lys	Trp	Xaa		Leu	
				95					100	4.4.4.				105		506
				gcc												586
Cys	Leu	HIS		Ala	ьеп	vai	ıyr		Asp	Pne	Pne	GIN	120	Pne	PIO	
			110					115		+~+	~	60 Y		226	tat	634
				ara Xaa												034
ъys	Add	125	261	Add	Wall	PHE	130	Deu	Буз	Cys	пец	135	110	7.511	*1*	
aad	cac		паа	gar	ata	act		aaa	aga	ata	cta		tta	aaa	ata	682
				Glu												
2,0	140	<i>د</i> رس	014	Ozu		145		-,-	• 5		150			-7-		
ata		agg	222	tgt	ttt		taσ	cact	ttc	aaac		ca c	ttta	taaai	t	733
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155		3	-1-	-7-	160											
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ccctttcatt gcaacttgca agtgagaaaa gatccttagt ggctctggtg gaagaaatag
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agtgaaaatg tttgaaacat gaaacatgtc tgtaggaagc atcagcatgg ccataagtgc
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                                                                      480
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                     -15
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Ser Lys Arg Ala Val Val Met Leu Lys Leu Glu Ile Thr Phe
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                                                                      932
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					tct Ser											251
					ttc Phe											299
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aaa	aaaa	aaa a	aa													363
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	1> p 2> 6		_	nal												
	1> p 2> 6			e												
	0> 3	-	tatt	tato	ct a	ttct	cttt	a ta	acto	catt	ctt	cttt	caa	tott	ctcagc	60
					t at	g ct	a ag u Ar	g at	a gc	c ct	t ac	a ct r Le	c at	C CC	a tct o Ser	112
					gct Ala 1											160
					tgc Cys	_		_								208
	-	_	_		cca Pro				araa	act	aatc	tcar	at t	ggca	gttaa	262
															caattg	322 382
				_	_			-							araaca cagttc	382 442
tcc	tcct	gtg	ctat	cttc	tc t	tcta	tcca	a gt	araa	tgta	ygc	cagg	arc	tcct	tecete	502 562
Lai	.caat		Lact	aadd	icy t	CCAA	guar	a at	yett	CCLL	Lta	caat	Cad	alla	ctgtat	202

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ggt gga aaa tac caa gtt ctt gga gat tac tct ttg gca gtg gtc ttc Gly Gly Lys Tyr Gln Val Leu Gly Asp Tyr Ser Leu Ala Val Val Phe -35 -20 -20	162
ccc ctg cac ttt tct gat cta att tct gtt tta tac ctt ata ccc aaa Pro Leu His Phe Ser Asp Leu Ile Ser Val Leu Tyr Leu Ile Pro Lys -15 -10 -5	210
aca ctt act acc aac aca gct gtt aaa cat tct ata caa aaa aat tgt Thr Leu Thr Thr Asn Thr Ala Val Lys His Ser Ile Gln Lys Asn Cys 1 5 10	258
atg mat ctg gta tta gga aaa tta ctt tca cag taaatatcaa agaaaaaaga Met Xaa Leu Val Leu Gly Lys Leu Leu Ser Gln 15 20	311
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gcc aga gcc ctg gac ggc tgc aga aat ggc att gcc cac cct gca agt Ala Arg Ala Leu Asp Gly Cys Arg Asn Gly Ile Ala His Pro Ala Ser .1 5 10	279									
gag aag cac aga ctc gag aaa tgt agg gaa ctc gag agc agc cac tcg Glu Lys His Arg Leu Glu Lys Cys Arg Glu Leu Glu Ser Ser His Ser 15 20 25	327									
gcc cca gga tca acc cag cac cga aga aaa aca acc aga aga	375									
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Cag cag cca ggt tcc ctc acc cca age tca ccc act gtt ggg gag att Gln Gln Pro Gly Ser Leu Thr Pro Ser Ser Pro Thr Val Gly Glu Ile -40 -35 -30	152									
atc tac aat aac acc aga aac aca ttg ggg tgg att ggg ggt atc ctt Ile Tyr Asn Asn Thr Arg Asn Thr Leu Gly Trp Ile Gly Gly Ile Leu -25 -20 -15 -10	200									
atg ggt tot ttt cag gga acc att gct gga caa ggc aca gga gcc acc Met Gly Ser Phe Gln Gly Thr Ile Ala Gly Gln Gly Thr Gly Ala Thr	248									

				-5					1				5			
		Ser					Gly					Pro		ggg Gly		296
999 61v	ctc	10 act	gtg Val	gcc Ala	cca	ccc	15 caa Gln	gcc Ala	gtc Val	agc Ser	ctc	20 cag	ggw	atc Ile	tac Tvr	344
_	25					30					35			rgg		392
														Xaa		
														tcc Ser 70		440
			${\tt Pro}$	rca				Leu	gaa				Asp	cct Pro		488
cga	sct	gtc	75 tgt	att	aat	CCC	cat	80 CCC	cca	cca	cca	atc	85 tta	aaa	abc	536
_		90	-				95					100		Lys		584
														Gly		
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gctt															cctgta	700
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- 55																
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	0> 3		רפתת	atcc	ta a	aatc	caaa	a ca	acca	agat	caa	aaca	act	cctc	aagagt	60
tac	tgat	cta	tnna ta a	tggc tg r	ag a wn n	gaaa nk t	aaaa tc a	a at ca g	tgtg ac c	acca cc t	gag: ct t	acgt ca g	gta tg a	gcaa at g	tgaaca aa aag lu Lys	120 171
220	acc	add		-	70				_	65					60	219
Lys	Arg	Arg	Glu -55	Arg	Glu	Glu	Arg	Gln -50	Asn	Ile	Val	Leu	Trp -45	Arg	Glņ	
cca	ctc	att	acc	tta	cad	tat	ttt	tct	cta	gaa	atc	ctt	qta	atc	ttq	267

ccg ctc att acc ttg cag tat ttt tct ctg gaa atc ctt gta atc ttg 267

Pro	Leu	Ile -40	Thr	Leu	Gln	Tyr	Phe	Ser	Leu	Glu	Ile	Leu -30	Val	Ile	Leu	
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													gtt Val			363
													ctt Leu 20			411
													ctt Leu			459
													gcc Ala			507
													gaa Glu			555
													gaa Glu			603
		_		_					_	~-		_	gcc Ala 100	_		651
													ttc Phe			699
													gag Glu			747
													gta Val			795
			_		_							_	agg Arg			843
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cccaagaaga ctgggga atg gag aga cag tca agg gtt atg tca gaa aag	170
Met Glu Arg Gln Ser Arg Val Met Ser Glu Lys	1,0
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gat gag tat cag ttt caa cat cag gga gcg gtg gag ctg ctt gtc ttc	218
Asp Glu Tyr Gln Phe Gln His Gln Gly Ala Val Glu Leu Leu Val Phe	220
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aat ttt ttg ctc atc ctt acc att ttg aca atc tgg tta ttt aaa aat	266
Asn Phe Leu Leu Ile Leu Thr Ile Leu Thr Ile Trp Leu Phe Lys Asn	
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cat cga ttc cgc ttc ttg cat gaa act gga gga gca atg gtg tat ggc	314
His Arg Phe Arg Phe Leu His Glu Thr Gly Gly Ala Met Val Tyr Gly	
10 15 20	
ctt aya atg gga cta att tta csa tat gct aca gca cca act gat att	362
Leu Xaa Met Gly Leu Ile Leu Xaa Tyr Ala Thr Ala Pro Thr Asp Ile	
25 30 35	
gaa agt ggr ret gte tat gae tgt gta aaa eta aet tte agt eea tea	410
Glu Ser Gly Xaa Val Tyr Asp Cys Val Lys Leu Thr Phe Ser Pro Ser	
40 45 50	
act ctg ctg gtt aat atc act gac caa gtt tat gar tat aaa tac aar	458
Thr Leu Leu Val Asn Ile Thr Asp Gln Val Tyr Glu Tyr Lys Tyr Lys	
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Arg Glu Ile Ser Gln His Xaa Ile Asn Pro His Xaa Gly Asn Ala Ile	
75 80 85	
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Pro Pro Ile Ile Phe His Ala Gly Tyr Ser Leu Lys Lys Arg His Phe	
105 110 115	
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Phe Gln Asn Leu Gly Ser Ile Leu Thr Tyr Ala Phe Leu Gly Thr Ala	
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ttt ctt ttg cta gaa ggc kaa aca gag caa gtr amn cat tca gag

Phe Leu Leu Glu Gly Gly Xaa Thr Glu Gln Val Xaa His Ser Glu -5 261 aca tat tgc atg ttt caa gac aag aag tac aga gtg ggt gag aga tgg Thr Tyr Cys Met Phe Gln Asp Lys Lys Tyr Arg Val Gly Glu Arg Trp 15 cat cct tac ctg gaa cct tat ggg ttg gtt tac tgc gtg aac tgc atc 309 His Pro Tyr Leu Glu Pro Tyr Gly Leu Val Tyr Cys Val Asn Cys Ile 30 357 tgc tca gag aat ggg aat gtg ctt tgc agc cga gtc aga tgt cca aat Cys Ser Glu Asn Gly Asn Val Leu Cys Ser Arg Val Arg Cys Pro Asn 50 40 405 gtt cat tgc ctt tct cct gtg cat att cct cat ctg tgc tgc cct cgc Val His Cys Leu Ser Pro Val His Ile Pro His Leu Cys Cys Pro Arg 65 60 450 tgc cca gaa gac tcc tta ccc cca gtg aac aat rwg gtg acc agc Cys Pro Glu Asp Ser Leu Pro Pro Val Asn Asn Xaa Val Thr Ser 80 tagtcttgck agtacaatgg gacaacttac caacatggas agctgttcgt agctgrrggg 510 570 ctctttcaga atcggcaacc cmatcaatgc acccagtgca gctgttcgga rggaaacktg 630 tattqtqqtc tcaaqacttq ccccaaatta acctgtgcct tcccagtctc tgttccarat 690 tectgetgee gggtwtgeag argagatgga caactgteat gggaacmtte tgatggtgat atcttccggc aacctgccaa cagagaagca agacattctt accaccgctc tcactatgat 750 810 cctccaccaa gccqacaggc tggaggtctg tcccgctttc ctggggccag aagtcaccgg 870 ggagctctta tggattccca gcaagcatca ggaaccattg tgcaaattgt catcaataac aaacacaagc atggacaagt gtgtgtttcc aatggaaaga cctattctca tggcgagtcc 930 990 tggcacccaa acctccgggc atttggcatt gtggagtgtg tgctatgtac ttgtaatgtc accaagcaag agtgtaagaa aatccactgc cccaatcgat acccctgcaa gtatcctcaa 1050 aaaatagacg gaaaatgctg caaggtgtgt ccaggtaaaa aagcaaaaaa aaaaaa 1106

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                                                                       167
agg att ctg cag tta atc ctg ctt gct ctg gca aca ggg ctt gta ggg
Arg Ile Leu Gln Leu Ile Leu Leu Ala Leu Ala Thr Gly Leu Val Gly
                                                 -5
                             -10
        -15
gga gag acc agg atc atc aag ggg ttc gag tgc aag cct cac tcc cag
                                                                       215
Gly Glu Thr Arg Ile Ile Lys Gly Phe Glu Cys Lys Pro His Ser Gln
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Pro Trp Gln Ala Ala Leu Phe Glu Lys Thr Arg Leu Leu Cys Gly Ala	
20 25 30	211
acg ctc atc gcc ccc aga tgg ctc ctg aca gcc gcc cac tgc ctc aag	311
Thr Leu Ile Ala Pro Arg Trp Leu Leu Thr Ala Ala His Cys Leu Lys 35 40 45	
	350
ccc cgc tac ata ktt cac ctg ggg cag cac aac ctc cag aag gag	359
Pro Arg Tyr Ile Xaa His Leu Gly Gln His Asn Leu Gln Lys Glu Glu 50 55 60	
	407
ggc tgt gag car acc cgg aca gcc act gag tcc ttc ccc cac ccc ggc Gly Cys Glu Gln Thr Arg Thr Ala Thr Glu Ser Phe Pro His Pro Gly	407
65 70 75	
ttc aac aac agc ctc ccc aac aaa gac cam mgc aat gac atc atg ctg	455
Phe Asn Asn Ser Leu Pro Asn Lys Asp Xaa Xaa Asn Asp Ile Met Leu	455
80 85 90 95	
gtg aak atg gma tcg cca gtc tcc atc acc tgg gct gtg cga ccc ctc	503
Val Xaa Met Xaa Ser Pro Val Ser Ile Thr Trp Ala Val Arg Pro Leu	503
100 105 110	
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acc ctc tcc tca cgc tgt gtc act gct ggc acc agc tgc ctc att tcc	551
Thr Leu Ser Ser Arg Cys Val Thr Ala Gly Thr Ser Cys Leu Ile Ser	
115 120 125 ggc tgg ggc agc acg tcc agc ccc cag tta cgc ctg cct cac acc ttg	E00
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130 135 140 cga tgc gcc aac atc acc atc att gag cac cag aag tgt gag aac gcc	647
Arg Cys Ala Asn Ile Thr Ile Ile Glu His Gln Lys Cys Glu Asn Ala	647
145 150 155	
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Asn Gln Ser Leu Gln Gly Ile Ile Ser Trp Gly Gln Asp Pro Cys Ala	731
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Ile Thr Arg Lys Pro Gly Val Tyr Thr Lys Val Cys Lys Tyr Val Asp	030
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Trp Ile Glu Thr Met Lys Asn Asn	000
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Ala Ala Val Ala	Pro Val Leu	Ser Ile Asn :	Ser Asp Phe Ser 5	Asp Leu
cgg gaa att aaa	aag caa ctg	ctg ctt att	cg ggc ctt acc	cgg gag 147
Arg Glu Ile Lys	15		20	
cgg ggc cta cta	cac agt agc	aaa tgg tcg	cg gag ttg gct	ttc tct 195
Arg Gly Leu Leu				Phe Ser
25	30		35	40
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Leu Pro Ala Leu	45	50		55
gag gaa gat gcc	cag gat atg	gat gcc tat	acc ctg gcc aag	g gcc tac 291
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60		65	70	
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Phe Asp Val Lys	GIU TYP Asp	Arg Ala Ala	ils Phe Led Al: 85	s Gly Cys
75 aat gca aga aaa	acc tot ttt	* -	· -	cta ata 387
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90	95	Dea 171	100	
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Arg Thr Asn Gly	Lys Val Lys			
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aggatggtaa aaaa				
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	-		1175													
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	_		1213													
\222	- 12	.05	121.	,				,								
-400	> 34	.4														
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															ly Leu	
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mg.	25	GIU	Mec	Ser	1111	30	1110		p	****	35			• • • •		
***		a+ a	tgc	200	aat	-	ata	<i>a</i> c=	a=a	999		200	taa	atc	cta	241
			Cys													231
	DET	vaı	Cyb	1111	45	пеп	ЦСИ	AIA	GIU	50	116	TAIL	110	vui	55	
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vai	TÀT	Arg	Thr	-	пур	TYL	пуъ	Arg		пуь	Ald	GIU	vaı	70	пуъ	
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			aaa													33/
GIN	ser	тÀг	Lys	Leu	GIU	гàг	ьys		GIU	inr	TTE	Thr		Ser	Ald	
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			cag													385
GIY	Arg		Gln	Lys	гÀг	гàг		GIU	Arg	хаа	хаа		хаа	ьеп	лаа	
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Asn		Asn	Arg	Asp	Leu		met	vaı	Arg	Met	-	Ser	met	Pne	Ala	
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Xaa	Gly	Leu	Ser	His	Arg	Asn	Leu		Gly	Asp	Asp	Thr		Asp	Cys	
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Ile	Gln	Lys	Ile	Leu	Gly	Leu	Ala	Pro	Ser	Arg	Ala	Ala	Thr	Lys	Gln	
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200	-	-			205					210	-					
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-90 -85	160
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-90 -85  aca atc gca aaa tyc rrg gcs tva gag ggc ctc cga gac ccc tat ggc Thr Ile Ala Lys Xaa Xaa Ala Xaa Glu Gly Leu Arg Asp Pro Tyr Gly -80 -75 -70  cgc ctc tgt ggt agc gag cac ccc cga aga cca cct gag cgg ccc gag Arg Leu Cys Gly Ser Glu His Pro Arg Arg Pro Pro Glu Arg Pro Glu -65 -60 -55  gaa gac ccg agc act cca gag gag gcc tct acc acc cct gaa gaa gcc Glu Asp Pro Ser Thr Pro Glu Glu Ala Ser Thr Thr Pro Glu Glu Ala -50 -45 -45 -40  tcg agc act gcc caa gca caa aag cct tca gtg ccc cgg agc aat ttt Ser Ser Thr Ala Gln Ala Gln Lys Pro Ser Val Pro Arg Ser Asn Phe -35 -30 -25 -20  cag ggc acc aag aaa agt ctc ctg atg tct ata tta gcg ctc atc ttc Gln Gly Thr Lys Lys Ser Leu Leu Met Ser Ile Leu Ala Leu Ile Phe -15 -10 -5	160 208 256 304 352
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tgg gat tcg gac gat gat gca gag gtt gag gct atc ctc aat tca ggt Trp Asp Ser Asp Asp Ala Glu Val Glu Ala Ile Leu Asn Ser Gly 95 100 105	688
gct arg ggt tat tcc gcc cct taagtaratc tgaggcagac ccttgggggt Ala Xaa Gly Tyr Ser Ala Pro 110 115	739
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Met Asn Val Gly Thr Ala His Xaa Xaa Val Asn Pro Asn Thr Arg	
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-20 -15 -10	
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Leu Leu His Ile Val Leu Leu Ser Ile Pro Phe Val Ser Val Pro Val	
-5 1 5 gtc tgg acc etc acc aac etc att cac aac atg gge atg tat ate tte	251
Val Trp Thr Leu Thr Asn Leu Ile His Asn Met Gly Met Tyr Ile Phe	231
10 15 20	
ctg cac acg gtg aag ggg aca ccc ttt gag acc ccg gac cag ggc aag	299
Leu His Thr Val Lys Gly Thr Pro Phe Glu Thr Pro Asp Gln Gly Lys 25 30 35 40	
25 30 35 40 gcg agg ctg cta acc cac tgg tgagcagatg gattatgggg tccagttcac	350
Ala Arg Leu Leu Thr His Trp	200
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	-

530

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ctggcattta gtctgcatta cacaaataga cactaattta tttggaacaa gcagcaaa
                                                                       298
atg aga act tta ttt ggt gca gtc agg gct cca ttt agt tcc ctc act
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Met Arg Thr Leu Phe Gly Ala Val Arg Ala Pro Phe Ser Ser Leu Thr
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 ctg ctt cta atc acc cct tct ccc agc cct ctt cta ttt gat aga ggt
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 ctg tee etc aga tea gea atg tet tageceetet ectetetee atteetteet
 Leu Ser Leu Arg Ser Ala Met Ser
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                                                                       508
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ggt ttg cgc ctt ttt ggc att ctt tca aca tgt cgt gta cat cac acc Gly Leu Arg Leu Phe Gly Ile Leu Ser Thr Cys Arg Val His His Thr	278
atg aat cag ttc cta att gat ata tct agc ttt acc tcc cga gtt aaa  Met Asn Gln Phe Leu Ile Asp Ile Ser Ser Phe Thr Ser Arg Val Lys  20 25 30	326
aaa aaa atc ttt tta ttt tat gcc ttc awa ggt tgc ycg ttt car agt Lys Lys Ile Phe Leu Phe Tyr Ala Phe Xaa Gly Cys Xaa Phe Gln Ser	374
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cct cts sta gct gag ccc act gca gag ggg gag cca cac ctg ccc acg Pro Leu Xaa Ala Glu Pro Thr Ala Glu Glv Glu Pro His Leu Pro Thr	158

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	-40					-35					-30						
Gly	cgg Arg	gas Xaa	byg Xaa	act Thr	Glu	gcc Ala	aac Asn	cgc Arg	ttc Phe	gcc Ala -15	tat Tyr	gct Ala	gcc Ala	ctc Leu	tgt Cys -10	206	
-25 ggc Glv	atc Ile	tcc Ser	ctg Leu	tcc Ser	-20 cag Gln	tta Leu	ttt Phe	cct Pro	gaa Glu	CCC	gaa Glu	cac His	agc Ser	tcc Ser	ttc	254	
tgc	aca	gag	ttc	-5 atg	gca	ggc	ctg	gtg	1 ckm	tgg	ctg	gag	5 ttg	tct	gaa	302	
-	Thr	10					15					20				350	
Ala	Val 25	Leu	Pro	Thr	Met	Thr 30	Ala	Phe	Ala	Ser	Gly 35	Leu	Gly	Gly	Glu		
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qqa	cga Arg	ccc Pro	gac Asp	Gly	gat Asp	cac His	tca Ser	gga Gly	cct Pro 65	tct Ser	gag Glu	ctt Leu	ctc Leu	act Thr 70	caa Gln	446	
	tgg Trp			60 tga	cscc	cgg (	gcca	gagt		cgtt	tgcc	a ca	tgac			498	
ctg	gaaa	aag tca	tgcc	cttg	ga g ga a	gagc tctg	tgga aaat	t gt g gc	cctt cgag	gaaa gcat	aga ccc	tgtt raaa	cct aaa	ggag aaaa	agcctg a	558 613	
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	21> p 22> 9	_		e													
	00> 3																
ac	cctat	caa	g at Me	g gt	c as	nc tt sn Ph	c co ne Pr	:o G]	ig aa ln Ly	aa at /s Il	t go	a gg a Gl	Ly Gl	u Le	tc tat u Tyr	50	
99 G1	a cct y Pro	Let	ato 1 Met	cto Lei	g gto 1 Val	tto Phe	? Thi	c te	g gtt ı Val	gct L Ala	ato a Ile -60	Let	a cto 1 Lei	cat His	Gly Gly	98	
at Me -5	g aag t Lys	acc	tct Ser	gad Asp	act Thi -50	c Ile	ato e Ile	c cgg	g gag g Glu	g ggq ı Gly	Thi	c cto	g ato 1 Met	g ggc Gly	aca Thr -40	146	
qc	c att	ggo Gly	c acc	c Cys	c tto s Pho	gge	tac Y Tyr	tgg r Tr	g ctg Lei -30	g gga u Gly	gto	t tca L Sex	tco r Ser	tto Phe	att E Ile	194	
+-				-3!		- t-a	- 22	- ac			- 200	ato	a cte		r ato	242	

tac ttc ctt gcc tac ctg tgc aac gcc cag atc acc atg ctg cag atg 242

Tyr Phe	Leu Ala -20	Tyr Leu	Cys A	Asn Ala -15	Gln	Ile	Thr	Met	Leu -10	Gln	Met	
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atc acc Ile Thr 10											_	338
ttg gtg Leu Val												386
cgg acc Arg Thr												434
gcc cta Ala Leu			Leu I									482
dtg gta Xaa Val 75		_	_	_								530
atc cag Ile Gln 90												578
ctt ccc Leu Pro												626
ctg cag Leu Gln		tgacccc	acc to	gaaattc	tt gg	JCC & S	gteet	ctt	tcc	egca		678
getgeaga tttgeage tgaaagge aaatgggt cccacatt aaaaaaat	aga ggar etg ccac eac aagg eca gctc egt cttg	tgagct g ccaaga a ctttga g	tageto ctccto aaccco	gcgt aa ggcc ag ctcc cc	gtaco gactg accta	tcc caa ccc	ttga ggct cttc	tgc ctgc	cag (	toggo ccaai cttta	cacttc tgcaga atctct	738 798 858 918 978 986

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tgg tgg tgc ttt cat ctt cag gca gaa gcc tct gcc cat ccc cct caa	344
Trp Trp Cys Phe His Leu Gln Ala Glu Ala Ser Ala His Pro Pro Gln -10 -5 1	
ggg ctg cag gcc caa ttc tca tgc tgc cct tgg gtg ggc atc tgt	389
Gly Leu Gln Ala Gln Phe Ser Cys Cys Pro Trp Val Gly Ile Cys	
5 10 15	
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Pro His Phe Leu Lys Ser Asn Leu Phe Leu Gln Leu Leu Val Ser His

-20 -15 -10 -5

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Glu Ile Val Cys Ala Thr Glu Thr Val Thr Thr Asn Phe Leu Arg His

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Leu Glu Gly Gly Ser Ala Arg His Val Val Phe Ser Gly Ser Gly Arg

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			His	cag			aag Lys	Leu	ckt				Thr	gag		350
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	_	45					50 tgg					55				446
Arg	Arg 60	His	Val	Pro	Ala	Xaa 65	Trp	Val	Leu	Leu	Xaa 70	Arg	Asp	Pro	Leu	440
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ggt				Ser	tcc		tgt Cys		Pro	cgt				Arg	gct	542
cra	rac	acc	cga	95 tcg	tgaa	aaac	tg o	tgas	100 sccas	jc ct	gtto	ctcc	g gg	105 cctra	aatg	597
Xaa	Xaa	Thr	Arg 110	Ser												
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atc	attt	tct													ctgtaa	60 120
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ggc					gtg Val		tgg Trp			cta					aac	265
ctg	cat	tgc	ttc			ctc	ccc	act	_	atg	cct	cta	ara	_	aaa	313

Leu His Cys Phe Pro Asp Leu Pro Thr Glu Met Pro Leu Xaa Ala Lys	
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Sly Xaa Asn Thr	
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gtttccaag ggaaggagca gcgtgtggga aagcacagaa gagtgagaag gaagcgacta lattttattt actttct atg cat cat ggc ctc aca cca ctg tta ctt ggt  Met His His Gly Leu Thr Pro Leu Leu Leu Gly  -50  -40  Ita cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa Ita His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys  -35  Ita cat gag ca tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt Ita Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu  -20  Ita cat gat gga tcg gca aga tat gga aga ctt cta ctt gag caa Ita Val Cys Cys Gly Ser Ala Ser Ile Val Ser Leu Leu Leu Glu Gln  -5  Iac att gat gta tct tct caa gat cta tct gga cag acg gcc aaa aag Isn Ile Asp Val Ser Ser Gln Asp Leu Ser Gly Gln Thr Ala Lys Lys  15  20  25  Iat gct gtt tct agt cgt cat aat gta att tgc cag tta ctt tct gac Iyr Ala Val Ser Ser Arg His Asn Val Ile Cys Gln Leu Leu Ser Asp  30  35  40  Iac aaa raa aaa cag atr cta aaa gtc tct tct gaa aac agc aat cca Iyr Lys Xaa Lys Gln Xaa Leu Lys Val Ser Ser Glu Asn Ser Asn Pro  45  50  50  60  60  60  60  60  60  60  6	110 158 206 254 302 350
determinant to the second seco	110 158 206 254 302 350 398

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Leu Val Leu Ala Ala Ala Gly Ala Val Ala Val Phe Leu Ile Leu														
-10 -5 1 5														
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Arg Ile Trp Val Val Leu Arg Ser Met Asp Val Thr Pro Arg Glu Ser														
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70					75					80	Mec	Add	1111	Буз	85		
tac	att	cac	cga	att		ara	age	caa	gaσ		can	cad	tcc	taa			393
Tyr	Ile	His	Ara	Ile	Pro	Xaa	Ser	Ara	Glu	Val	Gln	Cln	Ser	Trn	Dro		3,7,3
•			-	90				5	95		<b>Q.1.1.</b>	0111	501	100	110		
tcc	acc	gtt	tyc	acc	acc	ttg	cac	tcc	atq	tgg	ctc	tcc	ttk		cta		441
Ser	Thr	Val	Xaa	Thr	Thr	Leu	His	Ser	Met	Trp	Leu	Ser	Xaa	Pro	Leu		
			105					110					115				
att	cac	agg	gtg	aag	cca	rat	ttg	gtg	ttg	tgt	aac	gga	cca	gga	aca		489
Ile	His	Arg	Val	Lys	Pro	Xaa	Leu	Val	Leu	Cys	Asn	Gly	Pro	Gly	Thr		
		120					125					130					
tgt	gty	cct	atc	tgt	gta	tct	gcc	ctt	ctc	ctt	999	ata	cta	gga	ata		537
Суѕ		Pro	Ile	Cys	Val		Ala	Leu	Leu	Leu	Gly	Ile	Leu	Gly	Ile		
	135					140					145						
aag	aaa	gtg	atc	att	gtc	tac	gtt	gaa	agc	atc	tgc	cgt	gta	aaa	acs		585
	гÀЗ	vaı	11e	TIE		Tyr	Val	GIu	Ser		Cys	Arg	Val	Lys			
150	+				155			Am Am An		160					165		
Len	Cor	Met	CCC	gga	aag	TIO	ctg	חשה	cat	CEC	tca	aat	tac	ttc	att		633
пеп	Ser	Mec	Ser	170	гув	116	ьец	Pne	175	Leu	ser	Asn	Tyr		TTE		
att	cad	taa	cca		cta	222	722	220		~~~		+	gtg	180			<b>-01</b>
Val	Gln	Trn	Pro	Ala	Len	Lvs	Glu	Larg	Tur	Dro	Lvc	cox	Val	Tar	Tou		681
			185	,,,,,	Deu	<b></b>	014	190	TYL	FIU	шуъ	PET	195	IYI	neu		
aaa	cqa	att		tgad	aaat	aa d	aact		t ct	ttac	aatt	++0	cast	taa			733
Gly	Arg	Ile	Val			33		- 5			,		, cub c	·			,55
_	_	200															
cagt	arta	tg t	acto	aaat	t gg	9999	jaaaa	aaa	ccct	aca	tatt	tctt	gt a	aago	cgtct		793
gaca	gtcc	tg a	raat	tatt	gat	ggta	agga	ata	aaaa	atg	twca	gatr	ac t	cagt	gaara	l	853
aact	gagg	ict t	ctct	tate	ja aa	caaa	catt	gat	aaac	gta	acta	cyaa	at g	ıttta	tgcct		913
ctgt	aaac	ca a	attt	cttt	t ct	arat	aaaa	ata	tgta	tta	ctac	ctgo	aa a	aaaa	aaaaa	ı	973
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	·> 86																
	> DN																

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<222> 126..182

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<221> polyA site

<222> 856..867

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-15 -10 -5

			gcc													218
Gly	Gly	Phe	Ala	Ile 1	Val	Tyr	Tyr	Leu 5	Ile	Gln	Lys	Phe	His 10	Ser	Arg	
			tac													266
Thr	Leu	Tyr 15	Tyr	Lys	Leu	Ala	Val 20	Glu	Gln	Leu	Gln	Xaa 25	His	Pro	Glu	
			gct													314
Ala	Gln 30	Glu	Ala	Leu	Gly	Pro 35	Pro	Leu	Asn	Ile	His 40	Tyr	Leu	Lys	Leu	
			gaa													362
Ile	Asp	Arg	Glu	Asn	Phe	Val	Asp	Ile	Val		Ala	Lys	Leu	Lys		
45					50					55					60	
			gga													410
Pro	Val	Ser	Gly		Lys	Ser	Glu	Gly	Leu 70	Leu	Tyr	Val	His	Ser 75	Ser	
				65						~~~		<b>~</b> + ~			~~~	458
			CCC													430
_	_	_	Pro 80					85					90			
			ggt													506
Leu	Lys	Asp 95	Gly	Gln	Gln	Ile	Pro 100	Val	Phe	Lys	Leu	Ser 105	Gly	Glu	Asn	
ggt	gat	gaa	gtg	aaa	aag	gag	taga	agac	gac (	ccaga	aaga	cc c	agcti	tgcti	-	557
Gly	Asp	Glu	Val	Lys	Lys	Glu										
	110					115										
cta	gtcc	atc (	cttc	cctc	at c	tcta	ccata	a tg	gcca	ctgg	ggt	ggtg	gcc (	catc	ccagtg	617
aca	gaca	ctc	ctgc	aacc	ca gi	kttt	ccag	c ca	ccag	tggg	atg	atgg	tat 9	gtgc	cagcac	677
															ttaaac	737
tga	atcc	gaa a	agaa	actc	ct a	ttata	aaat	t taa	agat	aatg	taa	tgta	ttt	gaaa	gtgctt	797
tgt	ataa	aaa a	agca	catg	at a	aaag	gaat	c ag	aatt	aata	aaa	tgtt	tgt 1	tgate	ctttaa	857
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<222> 66..113
<223> Von Heijne matrix
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seq TALAAXTWLGVWG/VR

<210> 358

<221> polyA_signal <222> 490..495

<221> polyA_site <222> 508..519

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1 5 10 15

WO 99/31236 -274 - PCT/IB98/02122

aat gtc gac cgg ggc gcg ggc tcc atc cgg gaa gcc ggt ggg gcc ttc Asn Val Asp Arg Gly Ala Gly Ser Ile Arg Glu Ala Gly Gly Ala Phe	206
20 25 30 gga aag aga gag cag gct gaa gag gaa cga tat ttc cga gca cag agt Gly Lys Arg Glu Gln Ala Glu Glu Glu Arg Tyr Phe Arg Ala Gln Ser 35 40 45	254
aca gaa caa ctg gca rct ttg aaa aaa crc cat gaa gaa gar atc gtt Thr Glu Gln Leu Ala Xaa Leu Lys Lys Xaa His Glu Glu Glu Ile Val 50 55 60	302
cat cat aga gaa gga gat tgagcgtctg cagaaagaaa ttgagcgcca His His Arg Glu Gly Asp 65	350
taagcagaag atcaaaatgc tagaacatga tgattaagtg cacaccgtgt gccatagaat ggcacatgtc attgcccact tctgtgtaaa catggttctg gtttaactaa tatttgtctg tgtgctacta acagattata ataaattgtc atcagtgaaa aaaaaaaaa	410 470 519
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cac acc ttc att gtc ctg cac ctg gtc ttg caa ggg atg gtt tat act His Thr Phe Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr -15 -10 -5	159
gag tac acc tgg gaa gta ttt ggc tac tgt cag gag ctg gag ttg tcc Glu Tyr Thr Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser 1 5 10 15	207
ttg cat tac ctt ctt ctg ccc tat ctg ctg cta ggt gta aac ctg ttt Leu His Tyr Leu Leu Pro Tyr Leu Leu Gly Val Asn Leu Phe 20 25 30	255
ttt ttc acc ctg act tgt gga acc aat cct ggc att ata aca aaa gca Phe Phe Thr Leu Thr Cys Gly Thr Asn Pro Gly Ile Ile Thr Lys Ala 35 40 45	303
aat gaa tta tta ttt ctt cat gtt tat gaa ttt gat gaa ktg atg ttt Asn Glu Leu Leu Phe Leu His Val Tyr Glu Phe Asp Glu Xaa Met Phe 50 55 60	351
CCA aaa aac gtg agg tgc tct act tgt gat tta agg aaa cca gct cga Pro Lys Asn Val Arg Cys Ser Thr Cys Asp Leu Arg Lys Pro Ala Arg 65 70 75 80	399
tcc aas cac tgc akt gtg tgt aac tgg tgt gtg cac cgt ttc rac cat Ser Xaa His Cys Xaa Val Cys Asn Trp Cys Val His Arg Phe Xaa His	447

	tgt Cys	_					_			_						495
	ctc Leu			_	_			_	_	_	_	_		-	_	543
	gtg Val 130															591
	cag Gln															639
_	gtc Val				_		_		_						_	687
	atg Met	_		Phe	_	_	_	_				_				735
_	ttg Leu		-	_		_		_			_					783
	tac Tyr 210	-	-	-				_	_	_	_				•	831
	cct Pro	_		_	_			•								879
	ctt Leu						_					_			_	927
	gag Glu		_			_	tga	cmag	tgt a	atga	ctgc	et ti	gag	etgta	<b>a</b>	978
gtt	cccg	ttt :	attt	acac	at g	gga	tect	c gti	tttc	caaa	aaa	aaaa	aaa			1028

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 seq FACVPGASPTTLA/FP

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<221> polyA_site

<222> 441..452

<400> 360

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-55 -50 -45	158
aga tca tgc agc aga agc aga aaa agg caa acg aga agg agg	136
cca agt age tit gtg get teg tgt eca ace ete tig eee tite gee tgt	206
Pro Ser Ser Phe Val Ala Ser Cys Pro Thr Leu Leu Pro Phe Ala Cys -25 -20 -15	200
gtg cct gga gcc agt ccc acc acg ctc gcg ttt cct cct gta ktg ctc	254
Val Pro Gly Ala Ser Pro Thr Thr Leu Ala Phe Pro Pro Val Xaa Leu -10 -5 1 5	
aca ggt ccc avc acc gat ggc att ccc ttt gcc ctr nak tct gca gcg	302
Thr Gly Pro Xaa Thr Asp Gly Ile Pro Phe Ala Leu Xaa Ser Ala Ala 10 15 20	
ggt ccc ttt tgt gct tcc ttc ccc tca ggt avc ctc tct ccc cct ggg	350
Gly Pro Phe Cys Ala Ser Phe Pro Ser Gly Xaa Leu Ser Pro Pro Gly 25 30 35	
cca ctc ccg ggg gtg agg ggg tta ccc ctt ccc agt gtt ttt tat tcc	398
Pro Leu Pro Gly Val Arg Gly Leu Pro Leu Pro Ser Val Phe Tyr Ser 40 45 50	
tgt ggg gct cac ccc aaa gta tta aaa gta gct ttg taattcaaaa	444
Cys Gly Ala His Pro Lys Val Leu Lys Val Ala Leu 55 60 65	
aaaaaaaa	452
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C2137 HOMO Sapiens	
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cttagttcaa tggctgaaac acagtcgaaa agaatactgt gaattatgca agcacagatt	180
tgcttttaca ccaatttatt ctccagatat gccttcacgg cttccaattc aagacatatt	240
tgctggactg gttacaagta ttggcactgc aatacgatat tggtttcatt atacacttgt	300
ggcctttgca tggttgggag ttgttcctct tacagcatgt gagtattcat gcctctgatt	360 420
ggagttattt aaacattgca taactactta atattataaa gcaatattgc atcatattat	480
tatttgactg atgtttagtt atttgatgtc agagtgtcat gtattaggaa agccttactt	540
araaratgtt catcggaact aaraatgakt ttaacaggtc agttttttga gtgaatgtgg gaaaraacac agcatacaga atggctaacc atgaaagttc atgaaagcgt kgaaaaaatc	600
aaatcaaatc ataattagat atgaagt atg cta rag ctt tca agg gct aca aaa	654
Met Leu Xaa Leu Ser Arg Ala Thr Lys	
-25 -20	
rac ggc cgg gcg cgg tgg ctt atg cct gta atc cca gca ctt cag gag	702
Xaa Gly Arg Ala Arg Trp Leu Met Pro Val Ile Pro Ala Leu Gln Glu	
-15 -10 -5	
gcc gan gca ggc gga tca cga ggt cag gag ttt gaa act agc ctg gcc	750

Ala Xaa Ala Gly Gly Ser Arg Gly Gln Glu Phe Glu Thr Ser Leu Ala 1 5 10	
aac atg gag act gag gca gga gaa ttg ctt aaa ccc agg agg cgg agg Asn Met Glu Thr Glu Ala Gly Glu Leu Leu Lys Pro Arg Arg Arg 15 20 25	798
ttg car tgaactgaga tcgcaccact gcactccagc ttgggcaaca gagcaagact Leu Gln	854
30 · · ·	875
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cctcgagcg atg cac ctc ctt tcc aac tgg gca aac ccc gct tcc agc aga	111
Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg	
-10 -5 1	159
cgt cct tct atg gcc gct tca ggc act tct tgg ata tca tcg acc ctc Arg Pro Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu	137
5 10 15	
gca cac tot ttg tca ctg aga gac gtc tca gag agg ctg tgc agc tgc	207
Ala His Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys	
20 25 30	255
tgg agg act ata agc atg gga ccc tgc gcc cgg ggg tca cca atg aac Trp Arg Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn	233
35 40 45	
age tet gga gtg cac aga aaa tea age agg eta tte tac ate egg aca	303
Ser Ser Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr	
50 55 60 65	351
cca atg aga aga tct tca tgc cat tta gaa tgt crg gtt ata ttc ctt Pro Met Arg Arg Ser Ser Cys His Leu Glu Cys Xaa Val Ile Phe Leu	221
70 75 80	
ttg gga cgc caa ttg taaktgttac cttcaaagga tttccttttc taaaaaatta	406
Leu Gly Arg Gln Leu	
85	100
ttttaratgt ctaactttat gttattgctc acgggtattt gactgaattg ttgatttagg	466 526
ataagtcaat tootggaggg aaattaccaa ataaaatgat atgtatttot taccacaaaa aaaaa	531
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                                                                      111
          Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg
                      -10
eqt cet tet atg gee get tea gge act tet tgg ata tea teg ace etc
                                                                      159
Arg Pro Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu
                                10
                                                                      207
gca cac tot ttg toa ctg aga gac gto toa gag agg ctg tgc ago tgc
Ala His Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys
                            25
tgg agg act ata agc atg gga ccc tgc gcc cgg ggg tca cca atg aac
                                                                      255
Trp Arg Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn
    35
                        40
                                            45
                                                                      303
age tet gga gtg cae aga aaa tea age agg eta tte tae ate egg aca
Ser Ser Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr
                    55
                                        60
cca atg aga aga tct tca tgc cat tta raa tgt cag gtt ata ttc ctt
                                                                      351
Pro Met Arg Arg Ser Ser Cys His Leu Xaa Cys Gln Val Ile Phe Leu
                                    75
ttg gga cgc caa ttg tagteggtet tetettgeec aaccagacae tggcatecae
                                                                      406
Leu Gly Arg Gln Leu
tgtcttctgg cagtggctga accagagcca caatgcctgt gtcaactatg caaaccgcaa
                                                                      466
tgcraccaag ccttcacctg catccaagtt catccaggga tacctgggag ctgtcatcag
                                                                      526
cgccgtctcc attgctgtgg gccttatktc ctggttcaga aagccaacaa gttcacccca
                                                                      586
gecaceegee tteteateea gaggtttgtg cegtteeetg etgtageeag tgccaatate
                                                                      646
tgcaatgtgg tcctgatgcg gtacggggag ctggaggaag ggattgatgt cctggacagc
                                                                      706
                                                                      766
gatggcaacc tegtgggctc ctccaagate gcagecegac acgccctgct ggagacggcg
                                                                      826
ctgacgcgag tggtcctgcc catgcccatc ctggtgctac ccccgatcgt catgtccatg
                                                                      886
ctggagaaga cggctctcct gcaggcacgc ccccggctgc tcctccctgt gcaaagcctc
                                                                      946
gtgtgcctgg cagcettegg cctggccctg ccgctggcca tcagcetett cccgcaaatg
                                                                     1006
tragagattg aaacatccca attagagccg gagatagccc aggccacgag cagccggaca
gtggtgtaca acaaggggtt gtgagtgtgg tcagcggcct ggggacggag cactgtgcag
                                                                     1066
ccggggagct gaggggcarg gccgtagact cacggctgca cctgcaggga gcagcacgcc
                                                                     1126
aaccccagca gtcctgggcc ccctgggaga gtgctcaacc tacagtggag ggagactgac
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ccattcacat tttaacatag gcaagaggag ttctaacaca tttcgtacaa aaaaaaaa
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cocqtttgag ctcggtatcc tagtgcacac gccttgcaag cgacggcgcc atg agt
                                                                      116
                                                        Met Ser
                                                        -25
ctg act tcc agt tcc agc gta cga gtt gaa tgg atc gca gca gtt acc
                                                                      164
Leu Thr Ser Ser Ser Val Arg Val Glu Trp Ile Ala Ala Val Thr
            -20
                                 -15
                                                                      212
att get get ggg aca get gea att ggt tat eta get tae aaa aga ttt
Ile Ala Ala Gly Thr Ala Ala Ile Gly Tyr Leu Ala Tyr Lys Arg Phe
                                                                      260
tat gtt aaa gat cat cga aat aaa gct atg ata aac ctt cac atc cag
Tyr Val Lys Asp His Arg Asn Lys Ala Met Ile Asn Leu His Ile Gln
10
                    15
                                         20
aaa gac aac ccc aag ata gta cat gct ttt gac atg gag gat ttg gga
                                                                      308
Lys Asp Asn Pro Lys Ile Val His Ala Phe Asp Met Glu Asp Leu Gly
                30
                                     35
                                                                      356
gat aaa get gtg tac tgc cgt tgt tgg agg tcc aaa aag ttc cca ttc
Asp Lys Ala Val Tyr Cys Arg Cys Trp Arg Ser Lys Lys Phe Pro Phe
                                50
tgt gat ggg gct cac aca aaa cat aac gaa gag act gga gac aat gtg
                                                                      404
Cys Asp Gly Ala His Thr Lys His Asn Glu Glu Thr Gly Asp Asn Val
        60
                            65
ggc cct ctg atc atc aag aaa aaa gaa act taaatggaca cttttgatgc
                                                                      454
Gly Pro Leu Ile Ile Lys Lys Lys Glu Thr
                        80
tgcaaatcag cttgtcgtga agttacctga ttgtttaatt araatgacta ccacctctgt
                                                                      514
                                                                      574
ctgattcacc ttcgctggat tctaaatgtg gtatattgcm aactgcagct ttcacattta
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seq AMWLLCVALAVLA/WG
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score 8.39999961853027

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caatcagtgt gcctccacct tcaccatctt cttcccctta ctctcacttc cgtcatgtgt tttatacaac tctcaaatct ttcttggaga aggaggatat acatacataa tatgaaatgt gtttgttctt cacagtcacc cgattttact gatatttatt tgcattttac caataaaaag

757 781

<210> '366

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<213> Homo sapiens

aaaatgcaag ctcaaaaaaa aaaa

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<221> CDS

<222> 19..312

<221> sig_peptide

<222> 19..63

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<221> CDS

<222> 64..612

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 seq QLWLVMEFCGAGS/VT

<221> polyA_site

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gtt at Me	g gat t Asp														108
aac at Asn Me		_					His								156
ggt gc Gly Al -2	a Phe														204
ttg gt Leu Va -10				_		_			_		_	_			252
aac ac Asn Th															300
msg ga Xaa Gl															348
cat cg His Ar 40															396
gtt aa Val Ly 55															444
ggc ag Gly Ar															492
gtt at Val Il															540
gac tt Asp Le															588
ccc ct Pro Le	u Ser						tgag	gagci	tet (	cttc	ctcai	CC CC	ccgg	gaatc	642
cagege gettgg taegag caaaga	ctcg g taaa a acca a	aato accta	cacaq aatga	ge e	agaaq agcga gacaq	accaq ggtc	g caa	acaga	aaca	att	gatga	aag o	catco	cattta	702 762 822 849

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<213> Homo sapiens

<220>

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<222> 9..185

<221> sig_peptide

<222> 9..50

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<400> 369

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agg aac tgg ctg cca acc cct ccg gct acg ggc ccc tta ccg agc tcc Arg Asn Trp Leu Pro Thr Pro Pro Ala Thr Gly Pro Leu Pro Ser Ser 20 25 30	146
cag act ggt cat atg cgg atg gcc gcc ctg ctc ccc caa tgaaaggcca Gln Thr Gly His Met Arg Met Ala Ala Leu Leu Pro Gln 35 40 45	195
gcttcgaaaa aaagctgaaa gggagacktt tgcaaracra kttgtactgc tgtcacagga	255
aatggacgct ggattacaas catggcasct caggcagcar aakttgcagg aaraacaaag	315
gaagcaggaa aatgctctta aacccaaagg ggcttcactg aaaascccac ttccaaktca	375
ataaaaagca actcctgcct cccttcctca ccctgtctct ggatttcttt tctatcacct	435
aratgettea tecageeara aaatageett cackkteece atetgtette arageaaaar	495
agctgggacm ccaaraacaa gctgttarat cactgcctgg gaggcttggc ttartactct	555
catctctggt tocattocag ttcagctaag tottgcttta aaatttttac ctcctagctg	615
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coctactaaa gatacaaaca attagooggg cgtggtgggg tgcgcttgta atcccagcta	795
ctcaggaggc tgaggcagga gaatcgctta aactcgggag gtagaggttg cagtgagcca	855
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aaa	918
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tct cat gcc cgc ttt tat ttc tta ttt cat cga cca tta agg ctg tta	97
Ser His Ala Arg Phe Tyr Phe Leu Phe His Arg Pro Leu Arg Leu Leu	
-20 -15 -10	
aat ctg ctc atc ctt att gag ggc agt gtc gtc ttc tat cag ctc tat	145
Asn Leu Leu Ile Leu Ile Glu Gly Ser Val Val Phe Tyr Gln Leu Tyr	
-5 <b>1</b> 5	
tcc ttg ctg cgg tcg gag aag tgg aac cac aca ctt tcc atg gct ctc	193
Ser Leu Leu Arg Ser Glu Lys Trp Asn His Thr Leu Ser Met Ala Leu	
10 15 20	
atc ctc ttc tgc aac tac tat gtt tta ttt aaa ctt ctc cgg gac aga	241
Ile Leu Phe Cys Asn Tyr Tyr Val Leu Phe Lys Leu Leu Arg Asp Arg	
25 30 35 40	

40

25

		tta Leu														289
_	_	aac Asn		_	-				tgag	jggag	gaa c	teag	gataa	aa		336
attt	tgta		tatt	atgt											tttat	396 456 472
<211 <212	> 37 > 15 > DN > Ho	04	:apie	ens												
	> CI	s )10	92	,												
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	_	olyA_ 175														
	-	olyA_ 193	-													
	> 37															
		ag at	g cg	ja aa	ag gt	g gt	t tt	rat	t ac	ec gg	gg gc	t ag	gc ag	gt gg	gtgag gc att ly Ile	60 111
		gcc Ala	ctc				ctg	ctg								159
		ttg Leu								Ala						207
gct	ctq	cta								-15						
Ala				Ser					gag Glu 1	gtc					gtg Val	255
gat	Leu		Ala	Ser -5 ctg	His cag	Pro tca	Thr	Ala	Glu 1 cgg	gtc Val gcc	Thr	Ile aag	Val 5 gaa	Gln ctt	Val aag	255 303
gat Asp caa	Leu gtc Val agg	Leu agc Ser	Ala aac Asn cag	Ser -5 ctg Leu aga	His cag Gln tta	Pro tca Ser gac	Thr ttc Phe 15 tgt	Ala ttc Phe ata	Glu 1 cgg Arg	gtc Val gcc Ala	Thr tcc Ser aat	Ile aag Lys 20 gct	Val 5 gaa Glu 999	Gln ctt Leu atc	Val aag Lys atg	
gat Asp caa Gln cct	Leu gtc Val agg Arg 25 aat	agc Ser 10 ttt	Ala aac Asn cag Gln	Ser -5 ctg Leu aga Arg	His cag Gln tta Leu aat	Pro tca ser gac Asp 30 atc	Thr ttc Phe 15 tgt Cys	Ala ttc Phe ata Ile gca	Glu l cgg Arg tat Tyr	gtc Val gcc Ala cta Leu	Thr tcc ser aat Asn 35 ttt	lle aag Lys 20 gct Ala ggc	Val 5 gaa Glu 999 Gly	Gln ctt Leu atc Ile ttt	Val aag Lys atg Met tca	303
gat Asp caa Gln cct Pro 40 aga	Leu gtc Val agg Arg 25 aat Asn	agc ser 10 ttt Phe	Ala aac Asn cag Gln caa Gln	Ser -5 ctg Leu aga Arg cta Leu cat	His cag Gln tta Leu aat Asn 45 atg	Pro tca Ser gac Asp 30 atc Ile	Thr ttc Phe 15 tgt Cys aaa Lys tcc	Ala ttc Phe ata Ile gca Ala aca	Glu 1 cgg Arg tat Tyr ctt Leu	gtc Val gcc Ala cta Leu ttc Phe 50 gaa	Thr tcc Ser aat Asn 35 ttt Phe	aag Lys 20 gct Ala ggc Gly	Val 5 gaa Glu ggg Gly ctc Leu	Gln ctt Leu atc Ile ttt Phe	val aag Lys atg Met tca Ser 55 cag	303 351

gtc ttt ggc cat ttt atc ctg att cgg gaa ctg gag cct ctc ctc tgt Val Phe Gly His Phe Ile Leu Ile Arg Glu Leu Glu Pro Leu Leu Cys 90 95 100	543
cac agt gac aat cca tct cag ctc atc tgg aca tca tct cgc agt gca His Ser Asp Asn Pro Ser Gln Leu Ile Trp Thr Ser Ser Arg Ser Ala 105 110 115	591
agg aaa tot aat tot ago oto gag gac tot cag cac ago aaa ggc aag Arg Lys Ser Asn Phe Ser Leu Glu Asp Phe Gln His Ser Lys Gly Lys 120 125 130 135	639
gaa ccc tac agc tct tcc aaa tat gcc act gac ctt ttg agt gtg gct Glu Pro Tyr Ser Ser Ser Lys Tyr Ala Thr Asp Leu Leu Ser Val Ala 140 145 150	687
ttg aac agg aac ttc aac cag cag ggt ctc tat tcc aat gtg gcc tgt Leu Asn Arg Asn Phe Asn Gln Gln Gly Leu Tyr Ser Asn Val Ala Cys 155 160 165	735
cca ggt aca gca ttg acc aat ttg aca tat gga att ctg cct ccg ttt Pro Gly Thr Ala Leu Thr Asn Leu Thr Tyr Gly Ile Leu Pro Pro Phe 170 175 180	783
ata tgg acg ctg ttg atg ccg gca ata ttg cta ctt cgc ttt ttt gca  Ile Trp Thr Leu Leu Met Pro Ala Ile Leu Leu Leu Arg Phe Phe Ala  185 190 195	831
aat gca ttc act ttg aca cca tat aat gga aca gaa gct ctg gta tgg Asn Ala Phe Thr Leu Thr Pro Tyr Asn Gly Thr Glu Ala Leu Val Trp 200 205 210 215	879
ctt ttc cac caa aag cct gaa tct ctc aat cct ctg atc aaa tat ctg Leu Phe His Gln Lys Pro Glu Ser Leu Asn Pro Leu Ile Lys Tyr Leu 220 225 230	927
agt gcc acc act ggc ttt gga aga aat tac att atg acc cag aag atg Ser Ala Thr Thr Gly Phe Gly Arg Asn Tyr Ile Met Thr Gln Lys Met 235 240 245	975
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tatttaatat atatataaaa ccagagctga caatgacact ctggaacatt gcataccttc	
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<221> sig_peptide <222> 274..399

<223> Von Heijne matrix score 5.19999980926514

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tgaatgaaaa taaagctttg acttcaaaaa aagccagaat tgatccata atg gaa gaa	238												
Met Glu Glu													
-25													
ata agt tot coa ott gta gaa ttt gta aaa gtt ttg tgc acc aac cag	286												
Ile Ser Ser Pro Leu Val Glu Phe Val Lys Val Leu Cys Thr Asn Gln													
-20 -15 -10													
gtt ctc att act gcc agg gct gtg cct aca aaa aag gca tct gtg cga	334												
Val Leu Ile Thr Ala Arg Ala Val Pro Thr Lys Lys Ala Ser Val Arg													
-5 1 5													
tgt gtg gaa aaa agg ttt tgg ata cca aaa act aca agc aaa cat ctg	382												
Cys Val Glu Lys Arg Phe Trp Ile Pro Lys Thr Thr Ser Lys His Leu													
10 15 20 25													
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Ser Arg Cys Ile Asp Gly Ile Ser Gly Phe Leu Asn Asp Phe Thr Phe													
30 35 40													
tgc ctt gaa ttt tca agg cat aga tgt caa ctt aca gaa taacatgtkt	479												
Cys Leu Glu Phe Ser Arg His Arg Cys Gln Leu Thr Glu													
45 50													
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cttgctggtg tcatagttat tagaatcagc agcctcttaa ctaattgcgg tttcatagga	839												
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	gtg Val															254
vaı	vai	Tyr	5	пλа	PIO	Giu	GIII	10	ASII	UIS	MIG	PILE	15	TILL	Cys	
aat	gta	ttt	•	aca	ttg	gct	ttc		atq	ata	aat	gct		tcc	aat	302
	Val				_	_			_			_				
		20					25					30				
_	cag		_		_	_		-	_		_					350
Ala	Gln	Val	Arg	Gly	Asp		Tyr	Glu	Ser	Gly	_	Leu	Gly	Arg	Thr	
	35 gct		~++	+ ~ ~	c++	40	2++	cat	++~	a+~	45	ata		aaa	tca	398
	Ala															330
50	mu	**** 9	•		55			017		60				1	65	
ctt	att	gct	tcc	atg	tgg	att	ctt	ttt	ggt	gca	tat	gtt	acc	caa	aat	446
Leu	Ile	Ala	Ser	Met	Trp	Ile	Leu	Phe	Gly	Ala	Tyr	Val	Thr	Gln	Asn	
				70					75					80		
	gat															494
Thr	Asp	Val	Tyr 85	Pro	GIY	Leu	Ата	90	Pne	Pne	GIn	Asn	95	Leu	TTE	
+++	ttt	age		cta	atc	tac	222		gga	ада	acc	gaa		cta	taa	542
	Phe	_		_						_		_				
		100				•	105		2			110			•	
acc	tga	gatc	act	tctt	aagt	ca c	attt	tcct	t tt	gttai	tatt	ctg	tttg	tag		595
Thr																4==
					-		_		_		-	_			atgttt	655 715
_					_	_	-	_		-		_			tatttt tgagta	775
	-		-	_		_			-			-			catcat	835
	_			_	_	_		-			_	_		_	tgcctg	895
taa	tccc	agc .	actt	tggg.	ag g	ccga	ggcg	g gc	cgat	tgct	tga	ggtc	aag	tgtt	tgagac	955
															gcatgg	1015
															gaaccc	1075
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ctg cgc ttc ctg agg gct gac ggc gac ctg acg cta cta tgg gcc gag Leu Arg Phe Leu Arg Ala Asp Gly Asp Leu Thr Leu Leu Trp Ala Glu -5 10	149												
tgg cag gga cgc cca gaa tgg gag ctg act gat atg gtg gtg tgg Trp Gln Gly Arg Arg Pro Glu Trp Glu Leu Thr Asp Met Val Val Trp  15 20 25	197												
gtg act gga gcc tcg agt gga att ggt gag gag ctg gct tac cag ttg Val Thr Gly Ala Ser Ser Gly Ile Gly Glu Glu Leu Ala Tyr Gln Leu 30 35 40	245												
Ser Lys Leu Gly Val Ser Leu Val Leu Ser Ala Arg Arg Val His Glu 45 50 55	293												
ctg gaa agg gtg aaa aga aga tgc cta gag aat ggc aat tta aaa gaa Leu Glu Arg Val Lys Arg Arg Cys Leu Glu Asn Gly Asn Leu Lys Glu 60 65 70	341												
aaa gat ata ctt gtt ttg ccc ctt gac ctg acc gac act ggt tcc cat Lys Asp Ile Leu Val Leu Pro Leu Asp Leu Thr Asp Thr Gly Ser His 75 80 85 90	389												
gaa agc ggc tac caa agc tgt tct cca gga att tgg tagaatcgac Glu Ser Gly Tyr Gln Ser Cys Ser Pro Gly Ile Trp 95 100	435												
attotggtca acaatgtgga aatgtcccag cgttctctgt gcatggatac caacttggat	495												
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kctgcctcac atgatcgaga ngaarcaagg aaagattgtt actgtgaata gcatcctggg	615												
tatcatatct gtacctcttt ccattggata ctgtgctagc aagcatgctc tccggggktk	675												
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tggtaga	atc	gacat	tctg	g to	aaca	atgg	tgg	a at	g to	c ca	g cg	jt to	t ct	g tgc	175
JJ J			_	-		_		Me	et Se	er Gl	n Ar	g Se	r Le	eu Cys	
										-6	50				
atg gat	acc	agc	ttg	gat	gtc	tac	aga	rag	cta	ata	gag	ctt	aac	tac	223
Met Asp	Thr	Ser	Leu	Asp	Val	Tyr	Arg	Xaa	Leu	Ile	Glu	Leu	Asn	Tyr	
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tta ggg	acg	gtg	tcc	ttg	aca	aaa	tgt	gtt	ctg	cct	cac	atg	atc	gag	271
Leu Gly	Thr	Val	Ser	Leu	Thr	Lys	Cys	Val	Leu	Pro	His	Met	Ile	Glu	
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agg aag	g caa	gga	aag	att	gtt	act	gtg	aat	agc	atc	ctg	ggt	atc	ata	319
Arg Lys	Gln	Gly	Lys	Ile	Val	Thr	Val	Asn	Ser	Ile	Leu	Gly	Ile	Ile	
		-20					-15					-10			
tct gta	a cct	ctt	tcc	att	gga	tac	tgt	gct	agc	aag	cat	gct	ctc	cgg	367
Ser Val	l Pro	Leu	Ser	Ile	Gly	Tyr	Cys	Ala	Ser	Lys	His	Ala	Leu	Arg	
	-5					1				5					
ggt tt	ttt	aat	ggc	ctt	cga	aca	gaa	ctt	gcc	aca	tac	cca	ggt	ata	415
Gly Pho	e Phe	Asn	Gly	Leu	Arg	Thr	Glu	Leu	Ala	Thr	Tyr	Pro	Gly	Ile	
10			•	15					20					25	
ata gt	t tct	aac	att	tgc	cca	gga	cct	gtg	caa	tca	aat	att	gtg	gaa	463
Ile Va	l Ser	Asn	Ile	Cys	Pro	Gly	Pro	Val	Gln	Ser	Asn	Ile	Val	Glu	
			30	-				35					40		
aat to	c cta	gct	gga	gaa	gtc	aca	aaa	act	ata	ggc	aat	aat	gga	aac	511
Asn Se	r Leu	Ala	Gly	Glu	Val	Thr	Lys	Thr	Ile	Gly	Asn	Asn	Gly	Asn	
		45	-				50					55			
cag tc	c cac	aag	atg	aça	acc	agt	cgt	tgt	gtg	cgg	ctg	atg	tta	atc	559
Gln Se	r His	Lys	Met	Thr	Thr	Ser	Arg	Cys	Val	Arg	Leu	Met	Leu	Ile	
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agc at	g gcc	aat	gat	ttg	aaa	gaa	gtt	tgg	atc	tca	gaa	caa	cct	ttc	607
Ser Me	t Ala	Asn	Āsp	Leu	Lys	Glu	Val	Trp	Ile	Ser	Glu	Gln	Pro	Phe	
75				,	80					85					
ttg tt	a gta	aca	tat	ttg	tgg	caa	tac	atg	cca	acc	tgg	gcc	tgg	tgg	655
Leu Le	u Val	Thr	Tyr	Leu	Trp	Gln	Tyr	Met	Pro	Thr	Trp	Ala	Trp	Trp	
90				95					100					105	
ata ac	c aac	aag	atg	ggg	aag	aaa	agg	att	gag	aac	ttt	aag	agt	ggt	703
Ile Th	r Asr	Lys	Met	Gly	Lys	Lys	Arg	Ile	Glu	Asn	Phe	Lys	Ser	Gly	
		-	110					115					120		
gtg ga	t gen	n rac	tct	tct	tat	ttt	aaa	atc	ttt	aag	aca	aaa	cat	gac	751
Val As	p Āla	a Xaa	Ser	Ser	Tyr	Phe	Lys	Ile	Phe	Lys	Thr	Lys	His	Asp	
	•	125			_		130					135			
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caatct	tctt	atgo	ttct	ga a	taat	caaa	g ac	taat	ttgt	gat	ttta	ctt	ttta	atagat	871
atgact	ttgc	ttcc	aaca	tg o	rrtg	aaat	a aa	aaat	aaat	aat	aaaa	gat	tgcc	atgrrt	931
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 Pro
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 Ser
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 Gly
 Leu
 Cys
 Val
 Leu
 Gln
 Leu
 Thr
 Thr
 Thr
 -5

 Ala
 Val
 Thr
 Ser
 Ala
 Phe
 Leu
 Leu
 Ala
 Lys
 Val
 Asn
 Pro
 Phe
 Glu
 Xaa

 Phe
 Leu
 Ser
 Arg
 Gly
 Phe
 Trp
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 Ala
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 His

 Pro
 Cys
 Leu
 Asp
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 Ala
 Ala
 Ala
 His
 His

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<221> SIGNAL <222> -14..-1

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Trp Gly Thr Asp Gln Gly Ile Gly Gly Phe Gly Glu Glu Pro Gly Ile 20 25 30

Lys Ser Xaa Xaa Met Xaa Leu Ile Arg Ser Val Arg Thr Val Met Arg 35 40 45 50

Val Pro Leu Ile Ile Val Asn Ser Ile Ala Ile Val Leu Leu Leu 55 60

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                       -15
Arg Pro Ala Ser Ala Ala Pro Met Gly Gly Pro Glu Leu Ala Gln His
Glu Glu Leu Thr Leu Leu Phe His Gly Thr Leu Gln Leu Gly Gln Ala
                                20
Leu Asn Gly Val Tyr Arg Thr Thr Glu Gly Arg Leu Thr Lys Ala Arg
                            35
Asn Ser Leu Gly Leu Tyr Gly Arg Thr Ile Glu Leu Leu Gly Gln Glu
                        50
Val Ser Arg Gly Arg Asp Ala Ala Gln Glu Leu Arg Ala Ser Leu Leu
Glu Thr Gln Met Glu Glu Asp Ile Leu Xaa Leu Gln Ala Xaa Ala Thr
Ala Glu Val Leu Gly Glu Val Ala Gln Ala Gln Lys Val Leu Arg Asp
                                100
Ser Val Gln Arg Leu Xaa Xaa Gln Leu Xaa Xaa Ala Trp Leu Gly Pro
                                                120
                           115
Ala Tyr Arg Lys Phe Glu Val Leu Lys Ala Pro Pro Xaa Lys Gln Asn
                        130
                                           135
His Ile Leu Trp Ala Leu Thr Gly His Val Xaa Arg Gln Xaa Arg Glu
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<210> 381

<400> 382
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Met Val Ala Gln Gln Xaa Xaa Leu Xaa Gln Ile Gln Glu Lys Leu His

-50 -45 -55 Ser Gly Leu Ser Glu Val Val Glu Ala Ser Ser Leu Ser Trp Ser Thr -35 -30 Arg Ile Lys Gly Phe Ile Ala Cys Phe Ala Ile Gly Ile Leu Cys Ser -15 Leu Leu Gly Thr Val Leu Leu Trp Val Pro Arg Lys Gly Leu His Leu Phe Ala Val Phe Tyr Thr Phe Gly Asn Ile Ala Ser Ile Gly Ser Thr 15 Ile Phe Leu Met Gly Pro Val Lys Gln Leu Lys Arg Met Phe Glu Pro Thr Arg Leu Ile Ala Thr Ile Met Val Leu Leu Cys Phe Ala Leu Thr 50 Leu Cys Ser Ala Phe Trp Trp His Asn Lys Gly Leu Ala Leu Ile Phe 65 Cys Ile Leu Gln Ser Leu Ala Leu Thr Trp Tyr Ser Leu Ser Phe Ile 85 80 Pro Phe Ala Arg Asp Ala Val Lys Xaa Cys Phe Ala Val Cys Leu Ala 95 100

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<221> SIGNAL <222> -18..-1

<210> 384 <211> 64 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -22..-1

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Gln Lys Gly Ser Ala Met Glu Leu Ala Val Ile Thr Val Xaa Gly Val
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                       -5
Ala Lys Pro Asn Glu Gln Pro Trp Leu Leu Asn
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Leu Pro Thr Arg Gly Gln Thr Leu Lys Asp Thr Thr Ser Ser Ser
              1
-5
Ala Asp Ser Thr Ile Met Asp Ile Gln Val Pro Thr Arg Ala Pro Asp
       15 20 25 /
Ala Val Tyr Thr Glu Leu Gln Pro Thr Ser Pro Thr Pro Thr Trp Pro
    30 . 35
Ala Asp Glu Thr Pro Gln Pro Gln Thr Gln Thr Gln Gln Leu Glu Gly
                        55
Thr Asp Gly Pro Leu Val Thr Asp Pro Glu Thr His Xaa Ser Xaa Lys
                65
Ala Ala His Pro Thr Asp Asp Thr Thr Thr Leu Ser Glu Arg Pro Ser
                             85
Pro Ser Thr Xaa Val His Xaa Arg Pro Xaa Xaa Pro Ser Xaa His Leu
                         100 105
Val Phe Met Arg Met Thr Pro Ser Ser Met Met Asn Thr Pro Ser Gly
                      115 120
Asn Xaa Gly Cys Trp Ser Gln Leu Cys Cys Ser Ser Gln Ala Ser Ser
                   130
                                   135
Ser Ser Pro Val Ala Ser Ala Gly Ser Cys Pro Gly Tyr Ala Gly Ile
140 145 150
Ile Ala Gly Glu Ser Ile Arg Asn Arg Ser
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45 50 55 Leu Ala His Ala Val Ser Leu Thr Lys Leu Val Arg Gly Arg Lys Ala 60 65 70 Pro Phe Pro Val Gly Asp Ser Gly Ser Gly Arg Gly Leu Gln Pro Ser
75 80 85
Pro Gly Cys Tyr Arg Tyr
90 95

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<210> 390

WO 99/31236 -299- PCT/IB98/02122 .

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Asp Gly Cys Lys Ser Glu Ala Xaa Lys Phe Thr Val Arg Glu Ala Leu
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Lys Glu Asn Gln Val Ser Leu Thr Val Asn Arg Val Thr Ser Asn Asp
                               75
Ser Ala Ile Tyr Ile Cys Gly Ile Ala Phe Pro Ser Val Pro Glu Ala
                           90
Arg Ala Lys Gln Thr Gly Gly Gly Thr Thr Leu Val Val Arg Glu Ile
                       105
Lys Leu Leu Ser Lys Glu Leu Arg Ser Phe Leu Thr Ala Leu Val Ser
                   120
                                       125
Leu Leu Ser Val Tyr Val Thr Gly Val Cys Val Ala Phe Ile Leu Leu
               135
                                   140
Ser Lys Ser Lys Ser Asn Pro Leu Arg Asn Lys Glu Ile Lys Glu Asp
           150
                               155
Ser Gln Lys Lys Lys Ser Ala Arg Arg Ile Phe Gln Glu Ile Ala Gln
                           170
Glu Leu Tyr His Lys Arg His Val Glu Thr Asn Gln Gln Ser Glu Lys
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Asp Asn Asn Thr Tyr Glu Asn Arg Arg Val Leu Ser Asn Tyr Glu Arg
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35

Ser

<210> 395

<211> 73

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -24..-1

<400> 395

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Leu Trp Leu Gly Pro Ile Trp Pro Cys Ser Gly Ser Thr Leu Gly Lys
-5
1
5

Pro Asp Pro Gly Val Trp Pro Ser Leu Phe Arg Pro Trp Asp Ala Ala 10 15 20

Ser Pro Gly Asn Tyr Ala Leu Ser Arg Gly Xaa Asn Xaa Tyr Xaa Xaa 25 30 35 40

Trp Gly Gln Gly Thr His Ser Ser Leu

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<213> Homo sapiens

<220>

<221> SIGNAL

<222> -18..-1

<400> 396

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1 ' 5 10

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Gln Gly Thr Leu Thr Arg Leu Gln Ser Thr Pro Ala

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<211> 192

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<222> -93..-1

<400> 397

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-90 -85 -80

Tyr Met Arg Phe Ala Arg Ser Lys Arg Gly Leu Arg Leu Lys Thr Val

-70 Asp Ser Cys Phe Gln Asp Leu Lys Glu Ser Arg Leu Val Glu Asp Thr -50 -55 Phe Thr Ile Asp Glu Val Ser Glu Val Leu Asn Gly Leu Gln Ala Val -40 Val His Ser Glu Val Glu Ser Glu Leu Ile Asn Thr Ala Tyr Thr Asn -20 Val Leu Leu Leu Arg Gln Leu Phe Ala Gln Ala Glu Lys Trp Tyr Leu **-**5 Lys Leu Gln Thr Asp Ile Ser Glu Leu Glu Asn Arg Glu Leu Leu Glu 10 Gln Xaa Ala Glu Phe Glu Lys Ala Xaa Ile Thr Ser Ser Asn Lys Lys 25 Pro Ile Leu Xaa Val Thr Xaa Pro Lys Leu Ala Pro Leu Asn Glu Gly Gly Thr Ala Lys Leu Leu Asn Lys Val Ile Cys Ile Ile Leu Arg Asn 60 Gly Lys Ser Leu Ile Leu Ser Cys His Cys Leu Gly Trp Arg Asn Lys 75 Ser Gly Arg Phe Val Ser Gly Pro Leu Arg Ile Ile Ser Pro Leu Gln 90

<210> 398 <211> 149 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -72..-1

<400> 398

Met Asn Leu Phe Ile Met Tyr Met Ala Gly Asn Thr Ile Ser Ile Phe -65 Pro Thr Met Met Val Cys Met Met Ala Trp Arg Pro Ile Gln Ala Leu -45 -50 Met Ala Ile Ser Ala Thr Phe Lys Met Leu Glu Ser Ser Ser Gln Lys -35 -30 Phe Leu Gln Gly Leu Val Tyr Leu Ile Gly Asn Leu Met Gly Leu Ala -15 Leu Ala Val Tyr Lys Cys Gln Ser Met Gly Leu Leu Pro Thr His Ala 1 Ser Asp Trp Leu Ala Phe Ile Glu Pro Pro Glu Arg Met Glu Ser Val 15 Val Glu Asp Cys Phe Cys Glu His Glu Lys Ala Ala Pro Gly Pro Tyr 35 Val Phe Gly Ser Tyr Leu His Pro Ser Leu Ser Pro Val Ala Pro Gln 45 His Thr Leu Lys Leu Ile Thr Tyr Val Lys Lys Asn Gln Lys Thr Leu 65 Phe Ser Met Val Gly

<210> 399 <211> 73 <212> PRT <213> Homo sapiens

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<220>
<221> SIGNAL
<222> -20..-1
<400> 399
Met Thr Pro Leu Leu Thr Leu Ile Leu Val Val Leu Met Gly Leu Pro
                   -15
                                       -10
Leu Ala Gln Ala Leu Asp Cys His Val Cys Ala Tyr Asn Gly Asp Asn
               1
Cys Phe Asn Pro Met Arg Cys Pro Ala Met Val Ala Tyr Cys Met Thr
                           20
Thr Arg Thr Tyr Tyr Thr Pro Thr Arg Met Lys Val Ser Lys Ser Cys
                      35
Val Pro Arg Cys Phe Glu Xaa Cys Val
                  50
<210> 400
<211> 86
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -20..-1
<400> 400
Met Asn Leu His Phe Pro Gln Trp Phe Val His Ser Ser Ala Leu Gly
    -15
                                       -10
Leu Val Leu Ala Pro Pro Phe Ser Ser Pro Gly Thr Asp Pro Thr Phe
                               5
Pro Cys Ile Tyr Cys Arg Leu Leu Asn Met Ile Met Thr Arg Leu Ala
                           20
Phe Ser Phe Ile Thr Cys Leu Cys Pro Asn Leu Lys Glu Val Cys Leu
                       35
                                           40
Ile Leu Pro Glu Lys Asn Cys Asn Ser Arg His Ala Gly Phe Val Gly
                   50
                                       55
Pro Xaa Lys Leu Arg Gln
<210> 401
<211> 78
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -21..-1
<400> 401
Met Cys Pro Val Phe Ser Lys Gln Leu Leu Ala Cys Gly Ser Leu Leu
                        -15
                                           -10
Pro Gly Leu Trp Gln His Leu Thr Ala Asn His Trp Pro Pro Phe Ser
Xaa Phe Leu Cys Thr Val Cys Ser Gly Ser Ser Glu Gln Ile Ser Glu
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Tyr Thr Ala Ser Ala Thr Pro Pro Leu Cys Arg Ser Leu Asn Gln Glu
30 35 40
Pro Phe Val Ser Arg Ala Ile Arg Pro Lys Tyr Ser Ile Thr

50 45

<210> 402 <211> 65 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -28..-1 <400> 402 Met Gly Lys Gly His Gln Arg Pro Trp Trp Lys Val Leu Pro Leu Ser -25 Cys Phe Leu Val Ala Leu Ile Ile Trp Cys Tyr Leu Arg Glu Glu Ser

-5 -10 Glu Ala Asp Gln Trp Leu Arg Gln Val Trp Gly Glu Val Pro Glu Pro 10 15

-20

Ser Asp Arg Ser Glu Glu Pro Glu Thr Pro Ala Ala Tyr Arg Ala Arg 25

Thr

<210> 403 <211> 211 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -27..-1

<400> 403

Met Leu Leu Leu Ser Ile Thr Thr Ala Tyr Thr Gly Leu Glu Leu Thr -20

Phe Phe Ser Gly Val Tyr Gly Thr Cys Ile Gly Ala Thr Asn Lys Phe -5

Gly Ala Glu Glu Xaa Ser Leu Ile Gly Leu Ser Gly Ile Phe Ile Gly

Ile Gly Glu Ile Leu Gly Gly Ser Leu Phe Gly Leu Leu Ser Lys Asn 30

Asn Arg Phe Gly Arg Asn Pro Val Val Leu Leu Gly Ile Leu Val His 45

Phe Ile Ala Phe Tyr Leu Ile Phe Leu Asn Met Pro Gly Asp Ala Pro 60

Ile Ala Pro Val Lys Gly Thr Asp Ser Ser Ala Tyr Ile Lys Ser Ser 80

Lys Xaa Phe Ala Ile Leu Cys Xaa Phe Leu Xaa Gly Leu Gly Asn Ser 95 90

Cys Phe Asn Thr Xaa Leu Leu Xaa Ile Xaa Gly Phe Leu Tyr Ser Glu 110

Xaa Ser Ala Pro Xaa Phe Ala Ile Phe Asn Phe Val Gln Ser Ile Cys 125 130

Ala Ala Val Ala Phe Phe Tyr Ser Asn Tyr Leu Leu His Trp Gln 145 140

Leu Leu Val Met Val Ile Phe Gly Phe Xaa Gly Thr Ile Ser Phe Phe 160 155

Thr Val Glu Trp Glu Xaa Ala Ala Phe Val Xaa Arg Gly Ser Asp Tyr 175

Arg Ser Ile

<210> 404 <211> 123 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -80..-1 <400> 404 Met Ser Thr Trp Tyr Leu Ala Leu Asn Lys Ser Tyr Lys Asn Lys Asp -70 -75 Ser Val Arg Ile Tyr Leu Ser Leu Cys Thr Val Ser Ile Lys Phe Thr -60 -55 Tyr Phe His Asp Ile Gln Thr Asn Cys Leu Thr Thr Trp Lys His Ser -35 -45 -40 Arg Cys Arg Phe Tyr Trp Ala Phe Gly Gly Ser Ile Leu Gln His Ser -25 Val Asp Pro Leu Val Leu Phe Leu Ser Leu Ala Leu Leu Val Thr Pro -10 Thr Ser Thr Pro Ser Ala Lys Ile Gln Ser Leu Gln Ile Asp Leu Pro 10 Gly Gly Trp Arg Leu Ala Thr Asp Arg Ile Phe Thr Leu Ser Pro Val 25 Pro Met Asp Xaa Pro Leu Ile Leu His Gln Leu

<210> 405 <211> 86 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1 <400> 405 Met Glu Lys Ser Trp Met Leu Trp Asn Phe Val Glu Arg Trp Leu Ile -20 -15 Ala Leu Ala Ser Trp Ser Trp Ala Leu Cys Arg Ile Ser Leu Leu Pro -5 Leu Ile Val Thr Phe His Leu Tyr Gly Gly Ile Ile Leu Leu Leu Leu 15 10 Ile Phe Ile Ser Ile Xaa Gly Ile Leu Tyr Lys Phe Xaa Asp Val Leu 30 Leu Tyr Phe Pro Xaa Gln Xaa Ser Ser Ser Arg Leu Tyr Asp Ser His Ala His Trp Xaa Ser Xaa

<210> 406 <211> 162 <212> PRT <213> Homo sapiens

PCT/IB98/02122 -

<221> SIGNAL <222> -31..-1 <400> 406 Met Ala Ala Ala Trp Pro Ser Gly Pro Xaa Ala Pro Glu Ala Val Thr -25 -20 Ala Arg Leu Val Gly Val Leu Trp Phe Val Ser Val Thr Thr Gly Pro -10 ,<del>-</del> 5 Trp Gly Ala Val Ala Thr Ser Ala Gly Gly Glu Glu Ser Leu Lys Cys 10 Glu Asp Leu Lys Val Gly Gln Tyr Ile Cys Lys Asp Pro Lys Ile Asn 25 Asp Ala Thr Gln Glu Pro Val Asn Cys Thr Asn Tyr Thr Ala His Val 40 Ser Cys Phe Pro Ala Pro Asn Ile Thr Cys Lys Asp Ser Ser Gly Asn 55 Glu Thr His Phe Thr Gly Asn Glu Val Gly Phe Phe Lys Pro Ile Ser 75 70 Cys Arg Asn Val Asn Gly Tyr Ser Tyr Asn Glu Gln Ser His Val Ser 90 85 Phe Ser Trp Met Val Gly Ser Arg Ser Ile Leu Pro Trp Ile Pro Cys 105 110 Phe Gly Phe Val Lys Xaa Xaa His Cys Arg Val Xaa Trp Asn Trp Glu 125 Pro Asn 130

<210> 407 <211> 98 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -37..-1

<220>

<210> 408 <211> 70 <212> PRT <213> Homo sapiens

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<220>
<221> SIGNAL
<222> -15..-1
<400> 408
Met Arg Phe Leu Pro Cys Cys Leu Leu Trp Ser Val Phe Asn Pro Glu
                   -10
                                       +5
Ser Leu Asn Cys His Tyr Phe Xaa Xaa Glu Xaa Cys Ile Phe Xaa Ser
          5
Leu Gln Tyr Tyr Glu Ile Ser Leu Gln Glu Lys Leu Leu Gly Phe Leu
                          25
Trp Leu Cys Phe Leu Ser Tyr Phe Phe Arg Ala Val Tyr Phe Leu Ile
                       40
Asp Phe Ser Ser Phe Thr
50
<210> 409
<211> 60
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -45..-1
<400> 409
Met His Ser Leu Phe Ile Ala Ser Leu Lys Val Leu Phe Tyr Tyr Ser
                   -40
                                      -35
Phe Ser Phe Arg Phe Asn Trp Phe Asp Cys Leu Leu His Asn Leu Gly
               -25
                                   -20
                                                  -15
Glu Asn Phe Leu Ser Leu Leu Ser Lys Ser Cys Ser Ala Asp Pro Ser
         -10
                               -5
Gly Ser Thr Phe Met Arg Asp Ile Glu Thr Asn Lys
   5
                       10
<210> 410
<211> 39
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -22..-1
<400> 410
Met Pro Glu Ala Val Glu Gln Ser Ala His Leu Phe Val Thr Trp Ser
                          -15
                                       -10
Ser Gln Arg Ala Leu Ser His Pro Ala Pro Phe Leu Thr Xaa Xaa Lys
                                5
Asn Pro Phe Leu Trp Lys Leu
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<210> 411 <211> 51 <212> PRT

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<213> Homo sapiens
<220>
<221> SIGNAL
<222> -23..-1
<400> 411
Met Ala Phe Gln Ser Leu Leu Glu Met Lys Phe Phe Leu Cys Ala Ala
Phe Pro Leu Gly Ala Gly Val Lys Met Phe His Tyr Leu Gly Pro Gly
Lys Pro Leu Xaa Gln Ala Ser Pro Ser Pro His Pro His Arg Xaa Arg
10 15
Ile Trp Pro
<210> 412
<211> 95
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -48..-1
<400> 412
Met Ala Ser Ser His Trp Asn Glu Thr Thr Thr Ser Val Tyr Gln Tyr
                              -40
         -45
Leu Gly Phe Gln Val Gln Lys Ile Tyr Pro Phe His Asp Asn Trp Asn
                         -25
                                            -20
Thr Ala Cys Phe Val Ile Leu Leu Phe Ile Phe Thr Val Val Ser
                     -10
Leu Val Val Leu Ala Phe Leu Tyr Glu Val Leu Xaa Xaa Cys Cys
1 5
                       10
Val Lys Asn Lys Thr Val Lys Asp Leu Lys Ser Glu Pro Asn Pro Leu
                          25
Xaa Xaa Met Met Asp Asn Ile Arg Lys Arg Glu Thr Glu Val Val
                          40
<210> 413
<211> 60
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -32..-1
<400> 413
Met Asp Glu Tyr Ser Trp Trp Cys His Val Leu Glu Val Val Lys Gly
                          -25
Gln Met Phe Thr Phe Ile Asn Ile Thr Leu Trp Leu Gly Ser Leu Cys
                      -10
                                         - 5
Gln Arg Phe Phe Tyr Ala Ser Gly Thr Tyr Phe Leu Ile Tyr Ile Ser
                        10
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Thr Val Thr Pro Ser Trp Arg Leu Cys Leu Val Ser

<220>

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<210> 414
<211> 170
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -79..-1
<400> 414
Met Glu Asp Pro Asn Pro Glu Glu Asn Met Lys Gln Gln Asp Ser Pro
                -75
Lys Glu Arg Ser Pro Gln Ser Pro Gly Gly Asn Ile Cys His Leu Gly
           -60
                                -55
Ala Pro Lys Cys Thr Arg Cys Leu Ile Thr Phe Ala Asp Ser Lys Phe
                           -40
                                              -35
Gln Glu Arg His Met Lys Arg Glu His Pro Ala Asp Phe Val Ala Gln
                       -25
Lys Leu Gln Gly Val Leu Phe Ile Cys Phe Thr Cys Ala Arg Ser Phe
Pro Ser Ser Lys Ala Xaa Xaa Thr His Gln Arg Ser His Gly Pro Xaa
                               10
Ala Lys Pro Thr Leu Pro Val Ala Thr Thr Thr Ala Gln Pro Thr Phe
                           25
Pro Cys Pro Asp Cys Gly Lys Thr Phe Gly Gln Ala Val Ser Leu Xaa
                        40
Arg His Xaa Gln Xaa His Glu Val Arg Ala Pro Pro Gly Thr Phe Ala
                    55
                                       60
Cys Thr Xaa Cys Gly Gln Asp Phe Ala Gln Glu Xaa Gly Leu His Gln
               70
His Tyr Ile Arg His Ala Arg Gly Gly Leu
<210> 415
<211> 190
<212> PRT
<213> Homo sapiens
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<221> SIGNAL <222> -82..-1 <400> 415 Met Tyr Val Trp Pro Cys Ala Val Val Leu Ala Gln Tyr Leu Trp Phe -75 -70 His Arg Arg Ser Leu Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala Gly -60 Val Ser Leu Pro Gly Ile Leu Ala Ala Lys Cys Gly Ala Glu Val Ile -45 -40 Leu Ser Asp Ser Ser Glu Leu Pro His Cys Leu Glu Val Cys Arg Gln -30 -25 Ser Cys Gln Met Asn Asn Leu Pro His Leu Gln Val Val Gly Leu Thr -10 Trp Gly His Ile Ser Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp Ile 10 Ile Leu Ala Ser Asp Val Phe Phe Glu Pro Glu Xaa Phe Glu Asp Ile 20 25 Leu Ala Thr Ile Tyr Phe Leu Met His Lys Asn Pro Lys Val Gln Leu 35 40

<211> 114 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -60..-1 <400> 416 Met Met Ala Ala Val Pro Pro Gly Leu Glu Pro Trp Asn Arg Val Arg -55 -50 Ile Pro Lys Ala Gly Asn Arg Ser Ala Val Thr Val Gln Asn Pro Gly -40 -35 Ala Ala Leu Asp Leu Cys Ile Ala Ala Val Ile Lys Glu Cys His Leu -25 -20 -15 Val Ile Leu Ser Leu Lys Ser Gln Thr Leu Asp Ala Glu Thr Asp Val -5 Leu Cys Ala Val Leu Tyr Ser Asn His Asn Arg Met Gly Arg His Lys 10 Pro His Leu Ala Leu Lys Gln Val Glu Gln Cys Leu Lys Arg Leu Lys 30 Asn Met Asn Leu Glu Gly Ser Ile Gln Asp Leu Phe Glu Leu Phe Ser 45

<210> 417 <211> 161 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -108..-1

Ser Lys

<210> 416

<400> 417 Met Thr Ser Gly Gln Ala Arg Ala Ser Xaa Gln Ser Pro Gln Ala Leu -105 -100 Glu Asp Ser Gly Pro Val Asn Ile Ser Val Ser Ile Thr Leu Thr Leu -90 -85 -80 Asp Pro Leu Lys Pro Phe Gly Gly Tyr Ser Arg Asn Val Thr His Leu -70 -65 Tyr Ser Thr Ile Leu Gly His Gln Ile Gly Leu Ser Gly Arg Glu Ala -55 -50 His Glu Glu Ile Asn Ile Thr Phe Thr Leu Pro Thr Ala Trp Ser Ser -35 Asp Asp Cys Ala Leu His Gly His Cys Glu Gln Val Val Phe Thr Ala -20 Cys Met Thr Leu Thr Ala Ser Pro Gly Val Phe Pro Ser Leu Tyr Ser

<210> 418
<211> 67
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -21..-1

<210> 419 <211> 332 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -32..-1

<400> 419

Met Ile Xaa Leu Arg Asp Thr Ala Ala Ser Leu Arg Leu Glu Arg Asp -25 Thr Arg Gln Leu Pro Leu Leu Thr Ser Ala Leu His Gly Leu Gln Gln -10 - 5 Gln His Pro Ala Phe Ser Gly Val Ala Arg Leu Ala Lys Arg Trp Val Arg Ala Gln Leu Leu Gly Glu Gly Phe Ala Asp Glu Ser Leu Asp Leu Val Ala Ala Ala Leu Phe Leu His Pro Glu Pro Phe Thr Pro Pro Ser 40 Ser Pro Gln Val Gly Phe Leu Arg Phe Leu Phe Leu Val Ser Thr Phe 55 Asp Trp Lys Asn Asn Pro Leu Phe Val Asn Leu Asn Asn Glu Leu Thr 75 Val Glu Glu Gln Val Glu Ile Arg Ser Gly Phe Leu Ala Ala Arg Ala 90 Gln Leu Pro Val Met Val Ile Val Thr Pro Gln Xaa Arg Lys Asn Ser 100 105 110

```
Val Trp Thr Gln Asp Gly Pro Ser Ala Gln Ile Leu Gln Gln Leu Val
                        120
Val Leu Ala Ala Glu Xaa Leu Pro Met Leu Xaa Xaa Gln Leu Met Asp
                     135
                                      140
Pro Arg Gly Pro Gly Asp Ile Arg Thr Xaa Phe Arg Pro Pro Leu Asp
                  150
                                   155
Ile Tyr Asp Val Leu Ile Arg Leu Ser Pro Arg His Ile Pro Arg His
                  170
Arg Gln Ala Val Asp Ser Pro Ala Ala Ser Phe Cys Arg Gly Leu Leu
                            185
Ser Gln Pro Gly Pro Ser Ser Leu Met Pro Val Leu Gly Xaa Asp Pro
                        200
Pro Gln Leu Tyr Leu Thr Gln Leu Xaa Glu Ala Phe Gly Asp Leu Ala
                    215
                                       220
Leu Phe Phe Tyr Asp Gln His Gly Gly Glu Val Ile Gly Val Leu Trp
                230
                                   235
Lys Pro Thr Ser Phe Gln Pro Gln Pro Phe Lys Ala Ser Ser Thr Lys
                      250
             245
Gly Arg Met Val Met Ser Arg Gly Glu Leu Val Met Val Pro Asn
                           265
         260
Val Glu Ala Ile Leu Glu Asp Phe Ala Val Leu Gly Glu Gly Leu Val
      275 280 285
Gln Thr Val Glu Ala Arg Ser Glu Arg Trp Thr Val
                     295
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<210> 420
<211> 65
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -19..-1
<400> 420
Met Gly Gly Ile Trp Asn Ala Leu Ser Met Ser Ser Phe Ser Phe His
              -15
                   -10
Ser Ser Ser Cys Ser Ala Leu Ser Ala Lys Ser Leu Leu Ser Arg His
His Ile Leu Gln Gln Phe Leu Val Arg Lys Ser Val Pro Leu Glu Asn
                   20
Ala Ser Leu Pro Phe Pro His Leu Gly Ser Ser Leu Phe Lys Ile Val
30
              35
Gly
```

Thr Ser Met Met Leu Leu Thr Val Tyr Gly Gly Tyr Leu Cys Ser Val

-20

<210> 421
<211> 57
<212> PRT
<213> Homo sapiens

<220>
<221> SIGNAL
<222> -30..-1

<400> 421
Met Pro Thr Gly Lys Gln Leu Ala Asp Ile Gly Tyr Lys Thr Phe Ser

-10 -5 1

Arg Val Tyr His Tyr Phe Gln Trp Arg Arg Ala Gln Arg Gln Ala Ala
5 10 15

Glu Glu Gln Lys Xaa Ser Gly Ile Met
20 25

1 5 10 15

Asp Ser Asp Glu Leu Ala Ser Gly Xaa Phe Val Phe Pro Tyr Pro Tyr 20 25 30

Pro Phe Arg Pro Leu Pro Pro Ile Pro Phe Pro Arg Phe Pro Trp Phe 35 40 45

Arg Arg Asp Phe Pro Ile Pro Ile Pro Glu Ser Ala Pro Thr Thr Pro

Arg Arg Asn Phe Pro Ile Pro Ile Pro Glu Ser Ala Pro Inf Inf Pro
50 55 60

Leu Pro Ser Glu Lys 65

<210> 423 <211> 85 <212> PRT <213> Homo sapiens

<213> Homo sapiens

<220>
<221> SIGNAL
<222> -17..-1

<400> 423
Met Lys Lys Val Leu Leu Leu Ile Thr Ala Ile Leu Ala Val Ala Val
-15
-10
-5

Gly Phe Pro Val Ser Gln Asp Xaa Glu Arg Glu Lys Arg Ser Ile Ser

1 10 15

Asp Ser Asp Glu Leu Ala Ser Gly Phe Phe Val Phe Pro Tyr Pro Tyr 20 25 30

Pro Phe Arg Pro Leu Pro Pro Ile Pro Phe Pro Arg Phe Pro Trp Phe 35 40 45

Arg Arg Asn Phe Pro Ile Pro Ile Pro Glu Ser Ala Pro Thr Thr Pro 50 55 60

Leu Pro Ser Glu Lys 65

<210> 424 <211> 69 <212> PRT <213> Homo sapiens

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<220>
<221> SIGNAL
<222> -29..-1
<400> 424
Met Thr Cys Arg Gly Ser Cys Ser Tyr Ala Thr Arg Arg Ser Pro Ser
                                 -20
              -25
Glu Leu Ser Leu Leu Pro Ser Ser Leu Trp Val Leu Ala Thr Ser Ser
                           -5
          -10
Pro Thr Ile Thr Ile Ala Leu Ala Met Ala Ala Gly Asn Leu Cys Pro
                   10
                                        15
Leu Pro Ser Ser Xaa Arg Xaa Lys Arg Arg Trp Cys Gln Ala Xaa Gln
                                     30
Gln Xaa Ala Leu Leu
              40
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<210> 425 <211> 122

<212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -56..-1 <400> 425 Met Val Pro Trp Pro Arg Gly Lys Val Lys Thr Ala Pro Ile Pro Ile -50 Ser Arg Phe Pro Phe Leu Pro Thr His Asp Pro Pro Thr Pro Ala His -30 -35 Trp Ser Pro Ala Ser His Gln Gln Phe Lys His Xaa Ser Pro Leu Leu -15 Thr Leu Ala Leu Leu Gly Gln Cys Ser Leu Phe Xaa Asn Leu Arg Lys 1 Lys Leu Ala Gly Gln Lys Ala Lys Lys Leu Pro Ser Phe Ser Ser Leu 15 Pro Leu Thr Leu Trp Pro Leu Thr Pro Gln Phe Ala Glu Leu Thr Thr 35 30 Val Ala Gln Lys Lys Leu Arg Trp Ser Gly Thr Leu Gly Trp Gly Pro 45 Val Pro Ser Trp Val Gln Phe Phe Leu Gly 60

Arg Cys Ser Gly Ser Pro Leu Pro Leu 5 10

<210> 427
<211> 50
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -36..-1

<400> 427

Met Ala Pro His Thr Ala Ser Phe Gly Val Cys Pro Leu Leu Ser Val -35 -30 -25

Thr Arg Val Val Ala Thr Glu His Trp Leu Phe Leu Ala Ser Leu Ser -20 -15 -10 -5

Gly Ile Lys Thr Tyr Gln Ser Tyr Ile Ser Val Phe Cys Lys Val Thr 1 5 10

Leu Ile

<210> 428 <211> 136 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -18..-1

<400> 428 Met Asp Ser Leu Arg Lys Met Leu Ile Ser Val Ala Met Leu Gly Ala -15 -10 Xaa Ala Gly Val Gly Tyr Ala Leu Leu Val Ile Val Thr Pro Gly Glu Arg Arg Lys Gln Glu Met Leu Lys Glu Met Pro Leu Gln Asp Pro Arg 20 25 Ser Arg Glu Glu Ala Ala Arg Thr Gln Gln Leu Leu Leu Ala Thr Leu 40 Gln Glu Ala Ala Thr Thr Gln Glu Asn Val Ala Trp Arg Lys Asn Trp Met Val Gly Gly Glu Gly Gly Ala Thr Gly Xaa His Arg Glu Thr Gly 70 Leu Ala Ser Val Gly Ala Gly Pro Trp Leu Gly Arg Arg Asn Pro Arg 85 90 Gln Leu Ser Pro Ser Trp Ala Xaa Arg Lys Ile Arg Xaa Glu Asn Xaa 100 Met Pro Gly Leu Ser Gly Val Leu

<210> 429 <211> 194 <212> PRT <213> Homo sapiens

115 -

<221> SIGNAL <222> -65..-1

<400> 429 Met Gln Asp Ala Pro Leu Ser Cys Leu Ser Pro Thr Lys Trp Ser Ser -55 -60 Val Ser Ser Ala Asp Ser Thr Glu Lys Ser Ala Ser Ala Ala Gly Thr -45 -40 Arg Asn Leu Pro Phe Gln Phe Cys Leu Arg Gln Ala Leu Arg Met Lys -25 Ala Ala Gly Ile Leu Thr Leu Ile Gly Cys Leu Val Thr Gly Val Glu -10 Ser Lys Ile Tyr Thr Arg Cys Lys Leu Ala Lys Ile Phe Ser Arg Ala 10 Gly Leu Asp Asn Xaa Arg Gly Phe Ser Leu Gly Asn Trp Ile Cys Met 25 Ala Tyr Tyr Glu Ser Gly Tyr Asn Thr Thr Ala Gln Thr Val Leu Asp Asp Gly Ser Ile Asp Tyr Gly Ile Phe Gln Ile Asn Ser Phe Ala Trp 55 Cys Arg Arg Gly Lys Leu Lys Glu Asn Asn His Cys His Val Ala Cys 70 Ser Ala Leu Xaa Thr Asp Asp Leu Thr Asp Ala Ile Ile Cys Ala Xaa 90 Lys Ile Val Lys Glu Thr Gln Gly Met Asn Tyr Trp Gln Gly Trp Lys 100 105 Lys His Cys Glu Gly Arg Asp Leu Ser Xaa Trp Lys Lys Gly Cys Glu Val Ser

<210> 430 <211> 141 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -69..-1

<400> 430 Met Thr Ser Gln Pro Val Pro Asn Glu Thr Ile Ile Val Leu Pro Ser -65 Asn Val Ile Asn Phe Ser Gln Ala Glu Lys Pro Glu Pro Thr Asn Gln -45 Gly Gln Asp Ser Leu Lys Lys His Leu His Ala Glu Ile Lys Val Ile -30 Gly Thr Ile Gln Ile Leu Cys Gly Met Met Val Leu Ser Leu Gly Ile -15 Ile Leu Ala Ser Ala Ser Phe Ser Pro Asn Phe Thr Gln Val Thr Ser Thr Leu Leu Asn Ser Ala Tyr Pro Phe Ile Gly Pro Phe Phe Val Xaa 20 Lys Xaa Ser Glu Glu Gly Arg Met Gly Gln Xaa Gly Glu Glu Xaa Xaa 35 Asn Ser Leu Asn Phe Pro Xaa Ala Ser Leu Leu Xaa Leu Ile Cys Gln 50 Xaa Gln Gly Phe Asn Gly Glu Ser Cys Ser Pro Val Gly

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<210> 431
<211> 248
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -69..-1
<400> 431
Met Thr Ser Gln Pro Val Pro Asn Glu Thr Ile Ile Val Leu Pro Ser
               -65
                                    -60
Asn Val Ile Asn Phe Ser Gln Ala Glu Lys Pro Glu Pro Thr Asn Gln
                                -45
Gly Gln Asp Ser Leu Lys Lys His Leu His Ala Glu Xaa Lys Val Ile
       -35 -
                           -30
Gly Thr Ile Gln Ile Leu Cys Gly Met Met Val Leu Ser Leu Gly Ile
                       -15
Ile Leu Ala Ser Ala Ser Phe Ser Pro Asn Phe Thr Gln Val Thr Ser
Thr Leu Leu Asn Ser Ala Tyr Pro Phe Ile Gly Pro Phe Phe Phe Ile
                               20
Ile Ser Gly Ser Leu Ser Ile Ala Thr Lys Lys Arg Leu Thr Asn Leu
                           35
Leu Val His Thr Thr Leu Val Gly Ser Ile Leu Ser Ala Leu Ser Ala
                       50
Leu Val Gly Phe Ile Xaa Leu Ser Val Lys Gln Ala Thr Leu Asn Pro
                   65
                                       70
Ala Ser Leu Xaa Cys Glu Leu Xaa Lys Asn Asn Ile Pro Thr Xaa Xaa
                                   85
Tyr Val Xaa Tyr Phe Tyr His Asp Ser Leu Tyr Thr Thr Asp Xaa Tyr
                               100
Thr Ala Lys Ala Xaa Leu Ala Gly Thr Leu Ser Leu Met Leu Ile Cys
                           115
Thr Leu Leu Glu Phe Cys Xaa Xaa Val Leu Thr Ala Val Leu Arg Trp
                       130
                                           135
Lys Gln Ala Tyr Ser Asp Phe Pro Gly Ser Val Leu Phe Leu Pro Xaa
                   145
                                      150
Ser Tyr Ile Gly Asn Ser Gly Met Ser Ser Lys Met Thr His Asp Cys
               160
                                    165
Gly Tyr Glu Glu Leu Leu Thr Ser
            175
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Phe

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<210> 433
 <211> 86
 <212> PRT
 <213> Homo sapiens
 <220>
 <221> SIGNAL
 <222> -14..-1
 <400> 433
 Met Val Ala Leu Asn Leu Ile Leu Val Pro Cys Cys Ala Ala Trp Cys
                                   -5
                -10
 Asp Pro Arg Arg Ile His Ser Gln Asp Asp Val Leu Arg Ser Ser Ala
                                              15
                           10
 Ala Asp Thr Gly Ser Ala Met Gln Arg Arg Glu Ala Trp Ala Gly Trp
             25
. Arg Arg Ser Gln Pro Phe Ser Val Gly Leu Pro Ser Ala Glu Arg Leu
             40
                                    45
 Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg Ser Leu Val Gly Glu Gly
                                  60
 His Arg Ile Cys Asp Leu
             70
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<211> 144
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -58..-1
<400> 434
Met Thr Arg Leu Cys Leu Pro Arg Pro Glu Ala Arg Glu Asp Pro Ile
                               -50
Pro Val Pro Pro Arg Gly Leu Gly Ala Gly Glu Gly Ser Gly Ser Pro
                          -35
Val Arg Pro Pro Val Ser Thr Trp Gly Pro Ser Trp Ala Gln Leu Leu
                                          -15
                       -20
Asp Ser Val Leu Trp Leu Gly Ala Leu Gly Leu Thr Ile Gln Ala Val
                                      1
                   -5
Phe Ser Thr Thr Gly Pro Ala Leu Leu Leu Leu Leu Val Ser Phe Leu
          10
                               15
Thr Phe Asp Leu Leu His Arg Pro Ala Val Thr Leu Cys His Ser Ala
                          30
Asn Phe Ser Pro Gly Ala Arg Val Arg Gly Pro Val Lys Val Leu Asp
                       45
Ser Arg Arg Leu Tyr Ser Cys Lys Trp Val Gln Ser Gln Asp Asn Leu
                                       65
                   60
Ala Ser Arg Lys His Cys Cys Cys Cys Ser Trp Gly Trp Ala Arg Ser
```

<210> 434

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<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -16..-1
<400> 435
Met Glu Arg Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala
                        -10
Ser Ala Gly Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln
Cys Phe Lys Val Ser Ser Trp Thr Glu Cys Pro Pro Thr Trp Cys Ser
                                25
Pro Leu Asp Gln Val Cys Ile Ser Asn Glu Val Val Val Ser Phe Ser
                           40
Glu Ser Pro Pro Gly Arg Gly Xaa Val Pro Xaa Ala Gly Glu Xaa Pro
                        55
Val Pro Pro Pro Leu Xaa Asp Leu Xaa Met Thr Pro Arg Xaa Xaa Arg
                                        75
Ala Trp Gly Pro Val Gly Pro Lys Val Pro Pro Ala Val Ser Pro Ala
                                90
Leu Gly Ser Gly Glu His Pro Xaa Xaa
<210> 436
<211> 162
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -16..-1
<400> 436
Met Glu Arg Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala
                        -10
Ser Ala Gly Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln
                                   10
Cys Phe Lys Val Ser Ser Trp Thr Glu Cys Pro Pro Thr Trp Cys Ser
                               25
Pro Leu Asp Gln Val Cys Ile Ser Asn Glu Val Val Val Ser Phe Lys
Trp Ser Val Arg Val Leu Leu Ser Lys Arg Cys Ala Pro Arg Cys Pro
Asn Asp Asn Met Xaa Phe Glu Trp Ser Pro Ala Pro Met Val Gln Gly
                   70
                                        75
Val Ile Thr Arg Arg Cys Cys Ser Trp Ala Leu Cys Asn Arg Ala Leu
Thr Pro Gln Glu Gly Arg Trp Ala Leu Xaa Gly Gly Leu Leu Gln
                               105
Asp Pro Ser Arg Gly Xaa Lys Thr Trp Val Arg Pro Gln Leu Gly Leu
                           120
```

Pro Leu Cys Leu Pro Xaa Ser Asn Pro Leu Cys Pro Xaa Glu Thr Gln

140

135

Glu Gly 145

```
<210> 437
<211> 110
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -20..-1
<400> 437
Met Xaa Leu Met Val Leu Val Phe Thr Ile Gly Leu Thr Leu Leu Leu
                   -15
                                        -10
Gly Xaa Gln Ala Met Pro Ala Asn Arg Leu Ser Cys Tyr Arg Lys Ile
Leu Lys Asp His Asn Cys His Asn Leu Pro Glu Gly Val Ala Asp Leu
                            20
Thr Gln Ile Asp Val Asn Val Gln Asp His Phe Trp Asp Gly Lys Gly
                       35
                                           40
Cys Glu Met Ile Cys Tyr Cys Asn Phe Lys Arg Ile Ala Leu Leu Pro
                   50
                                       55
Lys Arg Arg Phe Leu Trp Thr Lys Asp Leu Phe Arg Asp Ser Leu Gln
                                   70
Gln Ser Met Arg Ile Phe Met Tyr Ser Gly Glu His His Ser
<210> 438
<211> 71
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -15..-1
<400> 438
Met Lys Leu Leu Thr His Asn Leu Leu Ser Ser His Val Arg Gly Val
                  -10
                                    -5
Gly Ser Arg Gly Phe Pro Leu Arg Leu Gln Ala Thr Glu Val Arg Ile
                               10
Cys Pro Val Glu Phe Asn Pro Asn Phe Val Ala Arg Met Ile Pro Lys
                           25
Val Glu Trp Ser Ala Phe Leu Glu Ala Xaa Asp Asn Leu Arg Leu Ile
Gln Val Pro Arg Arg Ala Gly
<210> 439
<211> 99
<212> PRT
<213> Homo sapiens
```

<221> SIGNAL <222> -24..-1 <400> 439 Met Lys Ser Ala Lys Leu Gly Phe Leu Leu Arg Phe Phe Ile Phe Cys -20 -15 -10

<220>

 Ser
 Leu
 Asn
 Thr
 Leu
 Leu
 Leu
 Gly
 Gly
 Val
 Asn
 Lys
 Ile
 Ala
 Glu
 Lys

 Ile
 Cys
 Gly
 Asp
 Leu
 Lys
 Asp
 Pro
 Cys
 Lys
 Leu
 Asp
 Met
 Asn
 Phe
 Gly

 Ser
 Cys
 Tyr
 Glu
 Val
 His
 Phe
 Arg
 Tyr
 Phe
 Tyr
 Asn
 Arg
 Thr
 Ser
 Lys

 Arg
 Cys
 Glu
 Thr
 Phe
 Val
 Phe
 Ser
 Cys
 Asn
 Gly
 Asn
 Leu
 Asn
 A

<210> 440 <211> 169 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -25..-1 <400> 440 Met Arg Lys Pro Ala Ala Gly Phe Leu Pro Ser Leu Leu Lys Val Leu -20 -15 Leu Leu Pro Leu Ala Pro Ala Ala Ala Gln Asp Ser Thr Gln Ala Ser Thr Pro Gly Ser Pro Leu Ser Pro Thr Glu Tyr Gln Arg Phe Phe Ala Leu Leu Thr Pro Thr Trp Lys Ala Glu Thr Thr Cys Arg Leu Arg Ala Thr His Gly Cys Arg Asn Pro Thr Leu Val Gln Leu Asp Gln Tyr Glu 45 50 Asn His Gly Leu Val Pro Asp Gly Ala Val Cys Ser Asn Leu Pro Tyr 65 Ala Ser Trp Phe Glu Ser Phe Cys Gln Phe Thr His Tyr Arg Cys Ser 80 Asn His Val Tyr Tyr Ala Lys Arg Val Leu Cys Ser Gln Pro Val Ser 95 Ile Leu Ser Pro Asn Thr Leu Lys Glu Ile Glu Xaa Ser Ala Glu Val 110 115 Ser Pro Thr Thr Asp Asp Leu Pro His Leu Thr Pro Leu His Ser Asp

130

125

Arg Thr Pro Asp Leu Pro Ala Leu Ala 140 Ala Asp Cys Gly Thr Ile Leu Leu Gln Asp Lys Gln Arg Lys Ile Tyr -50 -55 Cys Val Ala Cys Gln Glu Leu Asp Ser Asp Val Asp Lys Asp Asn Pro -35 Ala Leu Asn Ala Gln Ala Ala Leu Ser Gln Ala Arg Glu His Gln Leu -20 Ala Ser Ala Ser Glu Leu Pro Leu Gly Ser Arg Pro Ala Pro Gln Pro - 5 Pro Val Pro Arg Pro Glu His Cys Glu Gly Ala Ala Ala Gly Leu Lys Ala Ala Gln Gly Pro Pro Ala Pro Ala Val Pro Pro Asn Thr Xaa Val 30 25 Met Ala Cys Thr Gln Thr Ala Leu Leu Gln Lys Leu Thr Trp Ala Ser 45 Ala Glu Leu Gly Ser Xaa Thr Ser Xaa Gly Lys Xaa Ala Ser Ser Cys 60 Val Ala Leu Ser Ala His Val Arg Arg Pro Cys Ala Ala Cys Ser Ser Tyr Ser Thr Lys Arg Ser Pro

<210> 442 <211> 70 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1

<210> 443
<211> 381
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -33..-1

```
Lys Met Ala Thr Val Lys Ser Glu Leu Ile Glu Arg Phe Thr Ser Glu
                                      25
  Lys Pro Val His His Ser Lys Val Ser Ile Ile Gly Thr Gly Ser Val
             35
                                 40
  Gly Met Ala Cys Ala Ile Ser Ile Leu Leu Lys Gly Leu Ser Asp Glu
                             55
  Leu Ala Leu Val Asp Leu Asp Glu Xaa Lys Leu Lys Gly Glu Thr Met
                         70
  Asp Leu Gln His Gly Ser Pro Phe Thr Lys Met Pro Asn Ile Val Cys
                    85
                                         90
  Ser Lys Xaa Tyr Phe Val Thr Ala Asn Ser Asn Leu Val Ile Ile Thr
                                     105
  Ala Gly Ala Arg Gln Xaa Lys Gly Glu Thr Arg Leu Asn Leu Xaa Gln
             115
                                 120
  Arg Asn Val Ala Ile Phe Lys Leu Met Ile Ser Ser Ile Val Gln Tyr
  Ser Pro His Cys Lys Leu Ile Ile Val Ser Asn Pro Val Asp Ile Leu
                         150
                                             155
  Thr Tyr Val Ala Trp Lys Leu Ser Ala Phe Pro Lys Asn Arg Ile Ile
                     165
                                         170
. Gly Ser Gly Cys Asn Leu Ile Xaa Ala Arg Phe Arg Phe Leu Ile Gly
                 180
                                    185
  Gln Lys Leu Gly Ile His Ser Glu Ser Cys His Gly Trp Ile Leu Gly
                                 200
             195
  Glu His Gly Asp Ser Ser Val Pro Val Trp Ser Gly Val Asn Ile Ala
                            215
  Gly Val Pro Leu Lys Asp Leu Asn Ser Asp Ile Gly Thr Asp Lys Asp
                         230
                                             235
  Pro Glu Gln Trp Lys Asn Val His Lys Glu Val Thr Ala Thr Ala Tyr
                     245
                                         250
 Glu Ile Ile Lys Met Lys Gly Tyr Thr Ser Trp Ala Ile Gly Leu Ser
                 260
                                     265
 Val Ala Asp Leu Thr Glu Ser Ile Leu Lys Asn Leu Arg Arg Ile His
                                 280
  Pro Val Ser Thr Ile Thr Lys Gly Leu Tyr Gly Ile Xaa Glu Glu Val
                             295
                                                300
  Phe Leu Ser Ile Pro Cys Ile Leu Gly Glu Asn Gly Ile Thr Asn Leu
                         310
                                            315
  Ile Lys Ile Lys Leu Thr Pro Glu Glu Glu Ala His Leu Lys Lys Ser
                    325
                                         330
 Ala Lys Thr Leu Trp Glu Ile Gln Asn Lys Leu Lys Leu
                 340
```

-5

```
<210> 444
<211> 39
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -14..-1
<400> 444
Met Tyr Tyr Met Val Cys Leu Phe Phe Arg Leu Ile Phe Ser Glu His
                -10
Leu Pro Ile Ile Gly Thr Val Thr Ser His Lys Thr Gly Thr Leu Thr
```

Val Tyr Pro Thr Ser Ala Gly

```
<210> 445
<211> 50
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -37..-1
<400> 445
Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val Asn
                            -30
Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala Leu
                                           -10
                       -15
Ser Pro Cys Leu Thr Ala Pro Lys Ser Pro Arg Leu Ala Met Met Pro
                                5
-5
Asp Asn
<210> 446
<211> 51
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -26..-1
<400> 446
Met Thr Pro Trp Cys Leu Ala Cys Leu Gly Arg Arg Pro Leu Ala Ser
                        -20
                                            -15
Leu Gln Trp Ser Leu Thr Leu Ala Trp Cys Gly Ser Gly Ser His Trp
                    - 5
Thr Glu Arg Pro Xaa Gln Xaa Ser Pro Trp Xaa Ser Leu Ser Ala Thr
Thr Arg Gly
        25
<210> 447
<211> 242
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -30..-1
<400> 447
Met Gly Glu Ala Ser Pro Pro Ala Pro Ala Arg Arg His Leu Leu Val
                                        -20
                    -25
Leu Leu Leu Leu Ser Thr Leu Val Ile Pro Ser Ala Ala Ala Pro
                -10
                                    -5
 Ile His Asp Ala Asp Ala Gln Glu Ser Ser Leu Gly Leu Thr Gly Leu
                             10
 Gln Ser Leu Leu Gln Gly Phe Ser Arg Leu Phe Leu Lys Gly Asn Leu
```

25

Leu Arg Gly Ile Asp Ser Leu Phe Ser Ala Pro Met Asp Phe Arg Gly

40 Leu Pro Gly Asn Tyr His Lys Glu Glu Asn Gln Glu His Gln Leu Gly 60 Asn Asn Thr Leu Ser Ser His Leu Gln Ile Asp Lys Met Thr Asp Asn Lys Thr Gly Glu Val Leu Ile Ser Glu Asn Val Val Ala Ser Ile Gln 90 Pro Xaa Glu Gly Xaa Phe Glu Gly Asp Leu Lys Val Pro Arg Met Glu 105 110 Glu Lys Glu Ala Leu Val Pro Xaa Gln Lys Ala Thr Asp Ser Phe His 120 125 Thr Glu Leu His Pro Arg Val Ala Phe Trp Ile Ile Lys Leu Pro Arg 135 140 Arg Arg Ser His Gln Asp Ala Leu Glu Gly Gly His Trp Leu Xaa Glu 155 Lys Arg His Arg Leu Gln Ala Ile Arg Asp Gly Leu Arg Lys Gly Thr 170 His Lys Asp Xaa Leu Xaa Xaa Gly Thr Glu Ser Ser Ser His Ser Arg 185 190 Leu Ser Pro Arg Lys Xaa His Leu Leu Tyr Ile Leu Xaa Pro Ser Arg 200 205 Gln Leu

<210> 448
<211> 154
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -60..-1
<400> 448

Met Gly Ser Lys Cys Cys Lys Gly Gly Pro Asp Glu Asp Ala Val Glu -55 -50 Arg Gln Arg Arg Gln Lys Leu Leu Leu Ala Gln Leu His His Arg Lys -35 Arg Val Lys Ala Ala Gly Gln Ile Gln Ala Trp Trp Arg Gly Val Leu -25 -20 Val Arg Arg Thr Leu Leu Val Ala Ala Leu Arg Ala Trp Met Ile Gln -10 - 5 Cys Trp Trp Arg Thr Leu Val Gln Arg Arg Ile Arg Gln Arg Arg Gln 15 Ala Leu Leu Gly Val Tyr Val Ile Gln Glu Gln Ala Ala Val Lys Leu 30 Gln Ser Cys Ile Arg Met Trp Gln Cys Arg Gln Cys Tyr Arg Gln Met 45 Cys Asn Ala Leu Cys Leu Phe Gln Val Pro Lys Ser Ser Leu Ala Phe Gln Thr Asp Gly Phe Leu Gln Val Gln Tyr Ala Ile Pro Ser Lys Gln 75 Pro Glu Phe His Ile Glu Ile Leu Ser Ile 90

<210> 449 <211> 89 <212> PRT

<213> Homo sapiens

<220> <221> SIGNAL <222> -61..-1 <400> 449 Met Asn Ala Ala Ile Asn Thr Gly Pro Ala Pro Ala Val Thr Lys Thr -55 -50 Glu Thr Glu Val Gln Asn Pro Asp Val Leu Trp Asp Leu Asp Ile Pro -40 -35 Glu Ala Arg Ser His Ala Asp Gln Asp Ser Asn Pro Lys Ala Glu Ala -25 -20 Leu Leu Pro Cys Asn Leu His Cys Ser Trp Leu His Ser Ser Pro Arg -10 -5 Pro Asp Pro His Ser His Phe Pro Ser Xaa Arg Arg Cys Pro Leu Pro 10 His Pro Cys Ala Thr Tyr Pro Pro Xaa 25

<210> 450 <211> 73 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1

<210> 451 <211> 54 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -34..-1 <400> 451

Met Ile Pro Leu Ile Ser His Leu Ala Glu Ala Ala Pro Pro Thr Ser
-30 -25 -20

Trp Ser Leu Ile Ser Ser Val Leu Asn Val Gly His Leu Leu Phe Ser
-15 -10 -5

Ser Ala Cys Ser Val Ser Leu Glu Ala Leu Ser Thr Arg Asn Ile Lys
1 5 10

Ala Ile Ile Leu Met Lys
15

```
<210> 452
<211> 121
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -38..-1
<400> 452
Met Glu Ser Pro Gln Leu His Cys Ile Leu Asn Ser Asn Ser Val Ala
                                -30
Cys Ser Phe Ala Val Gly Ala Gly Phe Leu Ala Phe Leu Ser Cys Leu
                                            -10
                           -15
Ala Phe Leu Val Leu Asp Thr Gln Glu Thr Arg Ile Ala Gly Thr Arg
                       1
Phe Lys Thr Ala Phe Gln Leu Leu Asp Phe Ile Leu Ala Val Leu Trp
               15
                                  20
Ala Val Val Trp Phe Met Gly Phe Cys Phe Leu Ala Asn Gln Trp Gln
                               35
His Ser Pro Pro Lys Glu Xaa Leu Leu Gly Ser Ser Ser Ala Gln Ala
                           50
Ala Ile Gly Xaa His Leu Leu His Pro Cys Leu Asp Ile Pro Xaa
                       65
Leu Pro Gly Xaa Pro Gly Pro Pro Lys
<210> 453
<211> 166
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -37..-1
<400> 453
Met Ser Thr Val Gly Leu Phe His Phe Pro Thr Pro Leu Thr Arg Ile
                            -30
Cys Pro Ala Pro Trp Gly Leu Arg Leu Trp Glu Lys Leu Thr Leu Leu
                        -15
                                           -10
Ser Pro Gly Ile Ala Val Thr Pro Val Gln Met Ala Gly Lys Lys Asp
Tyr Pro Ala Leu Leu Ser Leu Asp Glu Asn Glu Leu Glu Glu Gln Phe
                                20
```

Gln Thr Arg Ser Val Asp Gln Asn Arg Lys Leu Ala Arg Lys Ile Leu 60 65 70 75

Gln Glu Lys Val Xaa Val Phe Tyr Asn Gly Glu Asn Ser Pro Val His 80 85 90

Lys Glu Lys Arg Glu Ala Ala Lys Lys Lys Gln Glu Arg Lys Lys Arg 95 100 105

Ala Lys Glu Thr Leu Glu Lys Lys Xaa Leu Leu Lys Xaa Leu Trp Glu

Val Lys Gly His Gly Pro Gly Gly Gln Ala Thr Asn Lys Thr Ser Asn

Cys Val Val Leu Lys Xaa Ile Pro Ser Gly Ile Val Val Lys Cys His

Ser Ser Lys Lys Val His 125

<210> 454 <211> 180 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1 <400> 454 Met Gly Ile Gln Thr Ser Pro Val Leu Leu Ala Ser Leu Gly Val Gly -25 -20 Leu Val Thr Leu Leu Gly Leu Ala Val Gly Ser Tyr Leu Val Arg Arg - 5 1 Ser Arg Arg Pro Gln Val Thr Leu Leu Asp Pro Asn Glu Lys Tyr Leu 15 Leu Arg Leu Leu Asp Lys Thr Thr Val Ser His Asn Thr Lys Arg Phe 30 Arg Phe Ala Leu Pro Thr Ala His His Thr Leu Gly Leu Pro Val Gly 45 Lys His Ile Tyr Leu Ser Thr Arg Ile Asp Gly Ser Leu Val Ile Arg 65 60 Pro Tyr Thr Pro Val Thr Ser Asp Glu Asp Gln Gly Tyr Val Asp Leu 80 Val Xaa Lys Val Tyr Leu Lys Gly Val His Pro Lys Phe Pro Glu Gly 95 Gly Lys Met Ser Xaa Tyr Leu Asp Xaa Leu Lys Val Gly Asp Xaa Val 110 Glu Phe Xaa Gly Pro Ser Gly Leu Leu Thr Tyr Thr Gly Lys Gly His 125 Phe Asn Ile Gln Pro Asn Lys Asn Leu His Gln Asn Pro Glu Trp Arg 145 140 Arg Asn Trp Glu

<210> 455 <211> 91 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -64..-1

<400> 455 Met Thr Pro Arg Ile Leu Ser Glu Val Gln Phe Ser Ala Phe Cys Pro -55 -60 Tyr Trp Thr Ile Ala Arg Ile Leu Glu Arg Val Gly Ser Ala Cys Phe -40 -45 Arg Leu Glu Leu Cys Ala Ala Ile Val Gly Tyr Phe Val Leu Asp Val -20 -25 Arg Thr Phe Leu Phe Ile Val Val Cys Val Ile Cys Val Thr Leu Asn - 5 -10 Phe Pro Arg Phe Tyr Phe Leu Cys Leu Ser Ser Leu Thr Ala Phe Gly Thr Pro Pro Ile Gly Val His Ile Pro Ser Pro

20

25

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<210> 456
<211> 257
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -23..-1
<400> 456
Met Arg Arg Ile Ser Leu Thr Ser Ser Pro Val Arg Leu Leu Xaa
            -20
                                -15
Leu Leu Leu Leu Ile Ala Leu Glu Ile Met Val Gly Gly His Ser
Leu Cys Phe Asn Phe Thr Ile Lys Ser Leu Ser Arg Pro Gly Gln Pro
Trp Cys Glu Ala His Val Phe Leu Asn Lys Asn Leu Phe Leu Gln Tyr
                                    35
Asn Ser Asp Asn Asn Met Val Lys Pro Leu Gly Leu Leu Gly Lys Lys
Val Tyr Ala Thr Ser Thr Trp Gly Glu Leu Thr Gln Thr Leu Gly Glu
                            65
Val Gly Arg Asp Leu Arg Met Leu Leu Cys Asp Ile Lys Pro Gln Ile
                        80
                                            85
Lys Thr Ser Asp Pro Ser Thr Leu Gln Val Xaa Xaa Phe Cys Gln Arg
                    95
                                        100
Glu Ala Glu Arg Cys Thr Gly Ala Ser Trp Gln Phe Ala Thr Asn Gly
                110
                                   115
Glu Lys Ser Leu Leu Phe Asp Ala Met Asn Met Thr Trp Thr Val Ile
                                130
Asn His Glu Ala Ser Xaa Ile Lys Glu Thr Trp Lys Lys Asp Arg Xaa
                            145
                                                150
Leu Glu Xaa Tyr Phe Arg Lys Leu Ser Lys Gly Asp Cys Asp His Trp
                        160
Leu Arg Glu Phe Leu Gly His Trp Glu Ala Met Pro Xaa Pro Xaa Val
                    175
                                        180
Ser Pro Xaa Asn Ala Ser Xaa Ile His Trp Ser Ser Ser Xaa Leu Pro
                190
                                    195
Xaa Xaa Trp Ile Ile Leu Gly Ala Phe Ile Leu Leu Xaa Leu Met Gly
           205
                               210
Ile Val Leu Ile Cys Val Trp Trp Gln Asn Gly Xaa Xaa Ser Thr Xaa
                            225
Хаа
```

```
<210> 457
<211> 193
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -60..-1
<400> 457
```

Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu Pro
-60 -55 -50 -45

Cys Ser Gly Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile Pro -35 -40 Leu Leu Cly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val Leu -20 -25 Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp Pro -5 Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala Pro 10 Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg Ala Val Gly Pro Thr Pro Gly Leu Leu Pro Glu Ala Ala Pro Xaa Thr 45 Xaa Gly Ala Leu Ser Ser Arg Ser Arg His Trp Ser Cys Ser Ile Val 60 Xaa Cys Leu His Leu His Xaa Leu Leu Ser Val Glu Thr Arg Xaa Phe 75 Xaa Lys His Leu Leu Val Leu Leu Val Ala Val Ala His Ser Val Leu 90 95 Glu Pro Pro Ala Leu Val Pro Asn Val Gln Cys Glu Met Cys Thr His 110 105 Ser Gly Pro Arg Asp Leu Glu Ala Ala Val Val Ser Pro Ala Pro Trp 125 Glu

<211> 107 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -28..-1 <400> 458 Met Val Leu Thr Leu Gly Glu Ser Trp Pro Val Leu Val Gly Arg Arg -20 -15 Phe Leu Ser Leu Ser Ala Ala Asp Gly Ser Asp Gly Ser His Asp Ser -5 Trp Asp Val Glu Arg Val Ala Glu Trp Pro Trp Leu Ser Gly Thr Ile 1.5 10 Arg Ala Val Ser His Thr Asp Val Thr Lys Lys Asp Leu Lys Val Cys 30 Val Glu Phe Xaa Gly Glu Ser Trp Arg Lys Arg Arg Trp Ile Glu Val 45 Tyr Ser Leu Leu Arg Lys Ala Phe Leu Val Lys His Asn Leu Val Leu 60 Ala Glu Arg Lys Ser Pro Glu Ile Ser Trp Gly

<210> 459 <211> 121 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -13..-1

<210> 458

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<400> 459
Met Leu Val Leu Arg Ser Ala Leu Thr Arg Ala Leu Ala Ser Arg Thr
            -10
                                -5
Leu Ala Pro Gln Met Cys Ser Ser Phe Ala Thr Gly Pro Arg Gln Tyr
Asp Gly Ile Phe Tyr Glu Phe Arg Ser Tyr Tyr Leu Lys Pro Ser Lys
                                        3.0
Met Asn Glu Phe Leu Glu Asn Phe Glu Lys Asn Ala Gln Leu Arg Thr
                                    45
Ala His Ser Glu Leu Val Gly Tyr Trp Ser Val Xaa Phe Gly Gly Arg
                                60
Met Xaa Thr Val Phe His Ile Trp Lys Tyr Asp Asn Phe Ala His Arg
                            75
Thr Glu Phe Gln Lys Ala Leu Ala Lys Asp Lys Glu Trp Gln Glu Gln
                        90
Phe Leu Ile Pro Asn Leu Ala Leu Asn
                   105
```

<210> 460
<211> 44
<212> PRT
<213> Homo sapiens

<220>
<221> SIGNAL
<222> -17..-1

<400> 460
Met Lys Val Gly Val Leu Trp Leu Ile Ser Phe Phe Thr Phe Thr Asp
-15
-10
-5

Gly His Gly Gly Phe Leu Gly Val Ser Trp Cys Tyr Val Ser Tyr Leu

1 5 10 15

Phe Ser Thr Asn Ser Pro Leu Ser Phe Arg Arg Ile
20 25

<210> 461 <211> 109 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -13..-1

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PCT/IB98/02122 --

85 90 95

<210> 462 <211> 143 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -41..-1 <400> 462 Met Ala Thr Ala Thr Glu Gln Trp Val Leu Val Glu Met Val Gln Ala -35 Leu Tyr Glu Ala Pro Ala Tyr His Leu Ile Leu Glu Gly Ile Leu Ile -15 Leu Trp Ile Ile Arg Leu Leu Phe Ser Lys Thr Tyr Lys Leu Gln Glu -5 Arg Ser Asp Leu Thr Val Lys Glu Lys Glu Glu Leu Ile Glu Glu Trp 15 Gln Pro Glu Pro Leu Val Pro Pro Val Pro Lys Asp His Pro Ala Leu Asn Tyr Asn Ile Val Ser Gly Pro Pro Ser His Lys Thr Val Val Asn 45 Gly Lys Glu Cys Ile Asn Phe Ala Ser Phe Asn Phe Leu Gly Leu Leu 65 Asp Asn Pro Arg Val Lys Ala Ala Ala Leu Ala Ser Leu Lys Lys Tyr 80 Gly Val Gly Thr Cys Gly Pro Cys Gly Phe Tyr Gly Thr Phe Glu

<210> 463

<211> 232

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -30..-1

<400> 463

Met Ala Ala Thr Ser Gly Thr Asp Glu Pro Val Ser Gly Glu Leu Val
-30 -25 -20 -15

Ser Val Ala His Ala Leu Ser Leu Pro Ala Glu Ser Tyr Gly Asn Xaa
-10 -5 1

Xaa Asp Ile Glu Met Ala Trp Ala Met Arg Ala Met Gln His Ala Glu
5 10 15

Val Tyr Tyr Lys Leu Ile Ser Ser Val Asp Pro Gln Phe Leu Lys Leu

20 25 30
Thr Lys Val Asp Asp Gln Ile Tyr Ser Glu Phe Arg Lys Asn Phe Glu

35 40 45 50 Thr Leu Arg Ile Asp Val Leu Xaa Pro Glu Xaa Leu Lys Ser Glu Ser

Ala Lys Glu Pro Pro Gly Tyr Asn Ser Leu Pro Leu Lys Leu Leu Gly
70 75 80

Thr Gly Lys Ala Ile Thr Lys Leu Phe Ile Ser Val Phe Arg Thr Lys 85 90 95

Lys Glu Arg Lys Glu Ser Thr Met Glu Glu Lys Lys Glu Leu Thr Val

```
105
Glu Lys Lys Arg Thr Pro Arg Met Glu Glu Arg Lys Glu Leu Ile Val
                120
                                       125
Glu Lys Lys Lys Arg Lys Glu Ser Thr Glu Lys Thr Lys Leu Thr Lys
               135
                                    140
Glu Glu Lys Lys Gly Lys Lys Leu Thr Lys Lys Ser Thr Lys Val Val
           150
                               155
Lys Lys Leu Cys Lys Val Tyr Arg Glu Gln His Ser Arg Ser Tyr Asp
                           170
Ser Ile Glu Thr Thr Ser Thr Thr Val Leu Leu Ala Gln Thr Pro Leu
                       185
Val Lys Cys Lys Phe Leu Tyr Asn
                   200
<210> 464
<211> 61
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -21..-1
<400> 464
Met Thr Phe Arg His Gln Asp Asn Ser Leu Met Phe Phe Ser Met Met
                       -15
Ala Thr Cys Thr Ser Asn Val Gly Phe Thr His Thr Thr Met Asn Cys
Ser Leu Thr Ser Pro Val Asp Phe Lys Asp Leu Leu Arg Val Leu Leu
                                20
Ile Lys Phe Gly Tyr Asp Arg Lys Ser Thr Ile Lys Ser
                            35
<210> 465
<211> 34
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -19..-1
<400> 465
Met Phe Leu Lys Ser Gly Ala Gly Leu Ser Ser Cys Leu Leu Pro Leu
                -15
                                    -10
Cys Trp Leu Glu Arg Lys Asp His Gly Arg Arg Pro Ser Xaa His Pro
                           5
Gly Arg
   15
<210> 466
<211> 215
<212> PRT
```

<220>

<213> Homo sapiens

```
<221> SIGNAL
<222> -54..-1
<400> 466
Met Asn Xaa Tyr Ala Ser Pro Phe Asn Xaa Gln Leu Xaa Tyr Leu Xaa
                               -45
             -50
Leu Ser Arg Phe Glu Cys Val His Arg Asp Gly Arg Val Ile Thr Leu
                            -30
Ser Tyr Gln Glu Gln Glu Leu Gln Asp Phe Leu Leu Ser Gln Met Ser
Gln His Gln Val His Ala Val Gln Gln Leu Ala Lys Val Met Gly Trp
 -5 1
Gln Val Leu Ser Phe Ser Asn His Val Gly Leu Gly Pro Ile Glu Ser
                               20
Xaa Gly Asn Ala Ser Ala Ile Thr Val Ala Pro Gln Val Val Thr Met
                  35
Leu Phe Gln Phe Val Met Asp Leu Lys Val Ala Ala Arg Leu Trp Phe
Ser Phe Leu Val Thr Asn Val Lys Thr Phe Gln Lys Val Met Phe Tyr
                    65
Lys Ile Thr Asn Gly Val Ile Phe Val Gly His Ser Lys Lys Phe Ser
                80
                                   85
Gly Ile Lys Trp Lys Val Xaa Ile Leu Phe Ile Lys Trp Xaa Cys Leu
                         100
Cys Leu His Leu Ala Leu Val Tyr Tyr Asp Phe Phe Gln Met Phe Pro
         110
                         115
Lys Xaa Val Ser Xaa Asn Phe Asp Leu Lys Cys Leu Gln Ile Asn Tyr
      125 130 135
Lys His Lys Glu Glu Ile Thr Ser Lys Arg Val Leu Phe Leu Lys Ile
 140 145
Ile Ile Arg Lys Cys Phe Ile
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<210> 468

<211> 85

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -24..-1

<400> 468
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        Met
        Cys
        Ser
        His
        Ala
        Ser
        Met
        Ser
        Phe
        His
        Thr
        Leu
        Phe
        His
        Leu
        Phe
        Leu
        Phe
        His
        Leu
        Leu
        Phe
        Phe</th
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Cys Leu Pro Cys Leu Ser Trp Asn Lys Lys Gly Asn Val Leu Gln
20 25 30

Pro Asn Phe
35

<210> 470 <211> 67 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -43..-1

<210> 471 <211> 63 <212> PRT <213> Homo sapiens

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<222> -15..-1
<400> 471
Met Gly Ile Leu Ser Thr Val Thr Ala Leu Thr Phe Ala Arg Ala Leu
        -10
                                      - 5
Asp Gly Cys Arg Asn Gly Ile Ala His Pro Ala Ser Glu Lys His Arg
                              10
Leu Glu Lys Cys Arg Glu Leu Glu Ser Ser His Ser Ala Pro Gly Ser
                         25
Thr Gln His Arg Arg Lys Thr Thr Arg Arg Asn Tyr Ser Ser Ala
<210> 472
<211> 179
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -58..-1
<400> 472
Met Ser Thr Gly Gln Leu Tyr Arg Met Glu Asp Ile Gly Arg Phe His
          -55
                              -50
Ser Gln Gln Pro Gly Ser Leu Thr Pro Ser Ser Pro Thr Val Gly Glu
                          -35
                                             -30
Ile Ile Tyr Asn Asn Thr Arg Asn Thr Leu Gly Trp Ile Gly Gly Ile
                      -20
                                        -15
Leu Met Gly Ser Phe Gln Gly Thr Ile Ala Gly Gln Gly Thr Gly Ala
                 - 5
                                    1
Thr Ser Ile Ser Glu Leu Cys Lys Gly Gln Glu Leu Glu Pro Ser Gly
                             15
Ala Gly Leu Thr Val Ala Pro Pro Gln Ala Val Ser Leu Gln Gly Ile
               30
Tyr Thr Leu Pro Trp Leu Leu Gln Leu Phe His Ser Thr Ala Leu Xaa
Xaa Xaa Gln Gln Pro Asn Gly Ser Leu Ser Leu Asn Ile Ser Ser Ser
                  60
                                      65
His Ala Pro Xaa Pro Xaa Thr Cys Thr Leu Glu Pro Gly Val Asp Pro
                                  80
Thr Arg Xaa Val Cys Ile Asn Pro His Pro Pro Pro Pro Ile Leu Lys
                   95
Xaa Pro Leu Ser Pro Tyr Pro Lys Pro Gln Leu Gly Thr His Ala Gly
Gln Val Asn
   120
<210> 473
<211> 238
<212> PRT
<213> Homo sapiens
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<220>

<221> SIGNAL <222> -71..-1

<400> 473 Met Xaa Xaa Phe Thr Asp Pro Ser Ser Val Asn Glu Lys Lys Arg Arg -65 Glu Arg Glu Glu Arg Gln Asn Ile Val Leu Trp Arg Gln Pro Leu Ile -50 -45 Thr Leu Gln Tyr Phe Ser Leu Glu Ile Leu Val Ile Leu Lys Glu Trp -35 -30 Thr Ser Lys Leu Trp His Arg Gln Ser Ile Val Val Ser Phe Leu Leu -15 Leu Leu Ala Gly Leu Ile Ala Thr Tyr Tyr Val Glu Gly Val His Gln Gln Tyr Val Gln Arg Ile Glu Lys Gln Phe Leu Leu Tyr Ala Tyr Trp Ile Gly Leu Gly Ile Leu Ser Ser Val Gly Leu Gly Thr Gly Leu His 35 Thr Phe Leu Leu Tyr Leu Gly Pro His Ile Ala Ser Val Thr Leu Ala 50 Ala Tyr Glu Cys Asn Ser Val Asn Phe Pro Glu Pro Pro Tyr Pro Asp 65 Gln Ile Ile Cys Pro Asp Glu Glu Gly Thr Glu Gly Thr Ile Ser Leu Trp Ser Ile Ile Ser Lys Val Arg Ile Glu Ala Cys Met Trp Gly Ile 95 100 Gly Thr Ala Ile Gly Glu Leu Pro Pro Tyr Phe Met Ala Arg Ala Ala 115 110 Arg Leu Ser Gly Ala Glu Pro Asp Asp Glu Glu Tyr Gln Glu Phe Glu 130 Glu Met Leu Glu His Ala Glu Ser Ala Gln Val Arg Thr Val Gly Ile 145 Glu Asn Arg Thr Leu Tyr Phe Phe Leu Lys Arg Leu Leu Arg 160

<210> 474 <211> 178 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -37..-1

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100
His Ala Gly Tyr Ser Leu Lys Lys Arg His Phe Phe Gln Asn Leu Gly
            115
                                  120
Ser Ile Leu Thr Tyr Ala Phe Leu Gly Thr Ala Ile Ser Cys Ile Val
 125
                      130
Ile Gly
140
<210> 475
<211> 96
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -21..-1
<400> 475
Met Ser Met Gln Phe Leu Phe Lys Met Val Ala Leu Cys Cys Cys Leu
                      -15
Trp Lys Ile Ser Gly Cys Glu Glu Val Pro Leu Thr Tyr Asn Leu Leu
                 1
                               5
Lys Cys Leu Leu Asp Lys Ala His Cys Val Leu Leu Thr Pro Cys Gly
                    20
Tyr Ile Phe Ser Leu Ile Ser Pro Glu Ile Leu Lys Leu Thr Leu Ile
Thr Leu Xaa Ile Leu Leu Ile Leu Lys Asn Leu His Leu Leu Trp Leu
                     50
Thr Val Ser Ser Xaa Cys Val His Arg Ser Ser Ala Arg Lys Glu Lys
                   65
<210> 476
<211> 41
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -24..-1
<400> 476
Met His Thr Phe Ala Asn Asp Arg Gly Leu Tyr Arg Ile Leu Leu
               -20
                                -15
His Phe Tyr Cys Leu Leu Arg Ser Ser Glu Tyr Ile Leu Gly Tyr Lys
Val Leu Gly Val Phe Phe Pro Ile Leu
                       15
<210> 477
<211> 113
<212> PRT
<213> Homo sapiens
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<221> SIGNAL <222> -27..-1

<210> 478 <211> 250 <212> PRT <213> Homo sapiens <220>

<221> SIGNAL <222> -18..-1

<400> 478

Met Arg Ile Leu Gln Leu Ile Leu Leu Ala Leu Ala Thr Gly Leu Val -10 Gly Glu Thr Arg Ile Ile Lys Gly Phe Glu Cys Lys Pro His Ser Gln Pro Trp Gln Ala Ala Leu Phe Glu Lys Thr Arg Leu Leu Cys Gly Ala Thr Leu Ile Ala Pro Arg Trp Leu Leu Thr Ala Ala His Cys Leu Lys Pro Arg Tyr Ile Xaa His Leu Gly Gln His Asn Leu Gln Lys Glu 55 Glu Gly Cys Glu Gln Thr Arg Thr Ala Thr Glu Ser Phe Pro His Pro 70 Gly Phe Asn Asn Ser Leu Pro Asn Lys Asp Xaa Xaa Asn Asp Ile Met 85 Leu Val Xaa Met Xaa Ser Pro Val Ser Ile Thr Trp Ala Val Arg Pro 100 105 Leu Thr Leu Ser Ser Arg Cys Val Thr Ala Gly Thr Ser Cys Leu Ile 115 120 Ser Gly Trp Gly Ser Thr Ser Ser Pro Gln Leu Arg Leu Pro His Thr 135 Leu Arg Cys Ala Asn Ile Thr Ile Ile Glu His Gln Lys Cys Glu Asn 150 155 Ala Tyr Pro Gly Asn Ile Thr Asp Thr Met Val Cys Ala Ser Val Gln 165 170 Glu Gly Gly Lys Asp Ser Cys Gln Gly Asp Ser Gly Gly Pro Leu Val 180 185 Cys Asn Gln Ser Leu Gln Gly Ile Ile Ser Trp Gly Gln Asp Pro Cys 200 Ala Ile Thr Arg Lys Pro Gly Val Tyr Thr Lys Val Cys Lys Tyr Val 215 220 210 Asp Trp Ile Gln Glu Thr Met Lys Asn Asn

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<210> 479

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<211> 151
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -21..-1
<400> 479
Met Ala Ala Ser Thr Ser Met Val Pro Val Ala Val Thr Ala Ala Val
          -15
Ala Pro Val Leu Ser Ile Asn Ser Asp Phe Ser Asp Leu Arg Glu Ile
Lys Lys Gln Leu Leu Ile Ala Gly Leu Thr Arg Glu Arg Gly Leu
                              20
Leu His Ser Ser Lys Trp Ser Ala Glu Leu Ala Phe Ser Leu Pro Ala
                           35
Leu Pro Leu Ala Glu Leu Gln Pro Pro Pro Pro Ile Thr Glu Glu Asp
                       50
Ala Gln Asp Met Asp Ala Tyr Thr Leu Ala Lys Ala Tyr Phe Asp Val
Lys Glu Tyr Asp Arg Ala Ala His Phe Leu His Gly Cys Asn Ala Arg
               80
                                  85
Lys Ala Tyr Phe Leu Tyr Met Tyr Ser Arg Tyr Leu Val Arg Ala Ile
                              100
           95
Leu Lys Cys His Ser Ala Phe Ser Glu Thr Ser Ile Phe Arg Thr Asn
                         115
      110
Gly Lys Val Lys Ser Phe Lys
  125
<210> 480
<211> 239
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -25..-1
<400> 480
Met Pro Arg Lys Arg Lys Cys Asp Leu Arg Ala Val Arg Val Gly Leu
                 -20
Leu Leu Gly Gly Gly Val Tyr Gly Ser Arg Phe Arg Phe Thr Phe
Pro Gly Cys Arg Ala Leu Ser Pro Trp Arg Val Arg Xaa Gln Arg Arg
                           15
Arg Cys Glu Met Ser Thr Met Phe Ala Asp Thr Leu Leu Ile Val Phe
                       30
Ile Ser Val Cys Thr Ala Leu Leu Ala Glu Gly Ile Thr Trp Val Leu
                   45
                                      50
Val Tyr Arg Thr Asp Lys Tyr Lys Arg Leu Lys Ala Glu Val Glu Lys
               60
                                   65
Gln Ser Lys Lys Leu Glu Lys Lys Lys Glu Thr Ile Thr Glu Ser Ala
           75
                              80
Gly Arg Gln Gln Lys Lys Ile Glu Arg Xaa Xaa Xaa Leu Xaa
```

Asn Asn Asn Arg Asp Leu Ser Met Val Arg Met Lys Ser Met Phe Ala 110 115 Ile Gly Phe Cys Phe Thr Ala Leu Met Gly Met Phe Asn Ser Ile Phe 125 130 Asp Gly Arg Val Val Ala Lys Leu Pro Phe Thr Pro Leu Ser Xaa Xaa 140 145 Xaa Gly Leu Ser His Arg Asn Leu Leu Gly Asp Asp Thr Thr Asp Cys 155 160 Ser Phe Ile Phe Leu Xaa Ile Leu Cys Thr Met Ser Ile Arg Gln Asn 170 175 Ile Gln Lys Ile Leu Gly Leu Ala Pro Ser Arg Ala Ala Thr Lys Gln 190 195 Ala Gly Gly Phe Leu Gly Pro Pro Pro Ser Gly Lys Phe Ser 205

<210> 481 <211> 208 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -92..-1

<400> 481

Met Arg Glu Pro Gln Lys Arg Thr Ala Thr Ile Ala Lys Xaa Xaa Ala -85 Xaa Glu Gly Leu Arg Asp Pro Tyr Gly Arg Leu Cys Gly Ser Glu His -70 Pro Arg Arg Pro Pro Glu Arg Pro Glu Glu Asp Pro Ser Thr Pro Glu -55 -50 Glu Ala Ser Thr Thr Pro Glu Glu Ala Ser Ser Thr Ala Gln Ala Gln -40 -35 Lys Pro Ser Val Pro Arg Ser Asn Phe Gln Gly Thr Lys Lys Ser Leu -20 Leu Met Ser Ile Leu Ala Leu Ile Phe Ile Met Gly Asn Ser Ala Lys Glu Ala Leu Val Trp Lys Val Leu Gly Lys Leu Gly Met Gln Pro Gly 10 15 Arg Xaa His Ser Ile Phe Gly Asp Pro Lys Lys Ile Val Thr Glu Xaa 25 30 Phe Val Arg Arg Gly Tyr Leu Ile Tyr Xaa Pro Val Pro Arg Xaa Ser 45 Pro Val Glu Tyr Xaa Phe Phe Trp Gly Pro Arg Ala His Val Glu Ser 60 Ser Xaa Leu Lys Xaa Xaa His Phe Val Ala Arg Val Arg Asn Arg Cys Ser Lys Asp Trp Pro Cys Asn Tyr Asp Trp Asp Ser Asp Asp Ala 90 95 Glu Val Glu Ala Ile Leu Asn Ser Gly Ala Xaa Gly Tyr Ser Ala Pro

<210> 482 <211> 86 <212> PRT <213> Homo sapiens <221> SIGNAL <222> -39..-1

<400> 482

Met Asn Val Gly Thr Ala His Xaa Xaa Val Asn Pro Asn Thr Arg Val -35 -30 -25

Met Asn Ser Arg Gly Ile Trp Leu Ser Tyr Val Leu Ala Ile Gly Leu
-20 -15 -10

Leu His Ile Val Leu Leu Ser Ile Pro Phe Val Ser Val Pro Val Val -5

Trp Thr Leu Thr Asn Leu Ile His Asn Met Gly Met Tyr Ile Phe Leu 10 20 25

His Thr Val Lys Gly Thr Pro Phe Glu Thr Pro Asp Gln Gly Lys Ala 30 35 40

Arg Leu Leu Thr His Trp
45

<210> 483

<211> 40

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -27..-1

<400> 483

Met Arg Thr Leu Phe Gly Ala Val Arg Ala Pro Phe Ser Ser Leu Thr
-25 -20 -15

Leu Leu Leu Ile Thr Pro Ser Pro Ser Pro Leu Leu Phe Asp Arg Gly
-10 -5 1 5

Leu Ser Leu Arg Ser Ala Met Ser

10

<210> 484

<211> 65

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -16..-1

<400> 484

Met Leu Gly Phe Phe Leu Phe Leu Ser Phe Val Leu Met Tyr Asp Gly
-15 -10 -5

Leu Arg Leu Phe Gly Ile Leu Ser Thr Cys Arg Val His His Thr Met 1 5 10 15

Asn Gln Phe Leu Ile Asp Ile Ser Ser Phe Thr Ser Arg Val Lys Lys 20 25 30

Lys Ile Phe Leu Phe Tyr Ala Phe Xaa Gly Cys Xaa Phe Gln Ser Ala 35 40 45

Thr

 Met Ala Met Trp Asn Arg Pro Xaa Xaa Xaa Leu Pro Gln Gln Pro Leu

 -55
 -50
 -45
 -40

 Xaa Ala Glu Pro Thr Ala Glu Gly Glu Pro His Leu Pro Thr Gly Arg
 -35
 -25

 Xaa Xaa Thr Glu Ala Asn Arg Phe Ala Tyr Ala Ala Leu Cys Gly Ile
 -20
 -15

 Ser Leu Ser Gln Leu Phe Pro Glu Pro Glu His Ser Ser Phe Cys Thr
 -5

 Glu Phe Met Ala Gly Leu Val Xaa Trp Leu Glu Leu Ser Glu Ala Val

 10
 15

 Leu Pro Thr Met Thr Ala Phe Ala Ser Gly Leu Gly Gly Glu Gly Xaa

30 35 40

Xaa Cys Val Cys Ser Asn Phe Thr Glu Gly Pro His Leu Glu Gly Arg
45 50 55

Pro Asn Gly Asn His Ser Gly Pro Ser Glu Leu Leu Thr Gln Gly Tro

Pro Asp Gly Asp His Ser Gly Pro Ser Glu Leu Leu Thr Gln Gly Trp
60 65 70

міа деі 75

<210> 486 <211> 209 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -84..-1

<400> 486 Met Val Asn Phe Pro Gln Lys Ile Ala Gly Glu Leu Tyr Gly Pro Leu -75 -80 Met Leu Val Phe Thr Leu Val Ala Ile Leu Leu His Gly Met Lys Thr -60 Ser Asp Thr Ile Ile Arg Glu Gly Thr Leu Met Gly Thr Ala Ile Gly -45 Thr Cys Phe Gly Tyr Trp Leu Gly Val Ser Ser Phe Ile Tyr Phe Leu -30 -25 Ala Tyr Leu Cys Asn Ala Gln Ile Thr Met Leu Gln Met Leu Ala Leu -15 -10 Leu Gly Tyr Gly Leu Phe Gly His Cys Ile Val Leu Phe Ile Thr Tyr 5 Asn Ile His Leu Arg Ala Leu Phe Tyr Leu Phe Trp Leu Leu Val Gly 20 Gly Leu Ser Thr Leu Arg Met Val Ala Val Leu Val Ser Arg Thr Val Gly Pro Thr Xaa Arg Xaa Leu Leu Cys Gly Thr Leu Ala Ala Leu His Met Leu Phe Leu Leu Tyr Leu His Phe Ala Tyr His Lys Xaa Val Xaa

Gly Ile Leu Asp Thr Leu Glu Gly Pro Asn Ile Pro Pro Ile Gln Arg 80 85 90 Val Pro Arg Asp Ile Pro Ala Met Leu Pro Ala Ala Arg Leu Pro Thr

```
100
       95
Thr Val Leu Asn Ala Thr Ala Lys Ala Val Ala Val Thr Leu Gln Ser
            115
His
125
<210> 487
<211> 36
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -17..-1
<400> 487
Met Gly Trp Gln Arg Trp Trp Cys Phe His Leu Gln Ala Glu Ala Ser
                   -10
Ala His Pro Pro Gln Gly Leu Gln Ala Gln Phe Ser Cys Cys Pro Trp
                  5
Val Gly Ile Cys
<210> 488
<211> 44
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -29..-1
<400> 488
Met Met Ser Ser Glu Leu Arg Arg Asn Pro His Phe Leu Lys Ser Asn
            -25 -20
Leu Phe Leu Gln Leu Leu Val Ser His Glu Ile Val Cys Ala Thr Glu
Thr Val Thr Thr Asn Phe Leu Arg His Glu Lys Ala
                       10
<210> 489
<211> 163
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -52..-1
<400> 489
Met Glu His Tyr Arg Lys Ala Gly Ser Val Glu Leu Pro Ala Pro Ser
       -50
                           -45
Pro Met Pro Gln Leu Pro Pro Asp Thr Leu Glu Met Arg Val Arg Asp
                       -30
```

Gly Ser Lys Ile Arg Asn Leu Leu Gly Leu Ala Leu Gly Arg Leu Glu

Gly Gly Ser Ala Arg His Val Val Phe Ser Gly Ser Gly Arg Ala Ala

-10

-15

-20

```
Gly Lys Ala Val Ser Cys Ala Glu Ile Val Lys Arg Arg Val Pro Gly
                           20
Leu His Gln Leu Thr Lys Leu Xaa Phe Leu Gln Thr Glu Asp Ser Trp
Val Pro Xaa Ser Pro Asp Thr Gly Leu Xaa Pro Leu Thr Val Arg Arg
                                        55
His Val Pro Ala Xaa Trp Val Leu Leu Xaa Arg Asp Pro Leu Asp Pro
                65
                                    70
Asn Glu Cys Gly Tyr Gln Pro Pro Gly Ala Pro Pro Gly Leu Gly Ser
                               85
Met Pro Ser Ser Cys Gly Pro Arg Ser Xaa Lys Arg Ala Xaa Xaa
                           100
Thr Arg Ser
   110
<210> 490
<211> 64
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -47..-1
<400> 490
Met His Gly Phe Glu Ile Ile Ser Leu Lys Glu Glu Ser Pro Leu Gly
                           -40
Lys Val Ser Gln Gly Pro Leu Phe Asn Val Thr Ser Gly Ser Ser Ser
                       -25
                                            -20
Pro Val Thr Trp Leu Gly Leu Leu Ser Phe Gln Asn Leu His Cys Phe
           -10
                                        -5
Pro Asp Leu Pro Thr Glu Met Pro Leu Xaa Ala Lys Gly Xaa Asn Thr
                                10
<210> 491
<211> 218
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -50..-1
<400> 491
Met His His Gly Leu Thr Pro Leu Leu Leu Gly Val His Glu Gln Lys
                    -45
                                        -40
Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys Ala Asn Leu Asn Ala
                -30
                                    -25
Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu Ala Val Cys Cys Gly
                                -10
            -15
Ser Ala Ser Ile Val Ser Leu Leu Glu Gln Asn Ile Asp Val Ser
Ser Gln Asp Leu Ser Gly Gln Thr Ala Lys Lys Tyr Ala Val Ser Ser
```

25

40

Arg His Asn Val Ile Cys Gln Leu Leu Ser Asp Tyr Lys Xaa Lys Gln

Xaa Leu Lys Val Ser Ser Glu Asn Ser Asn Pro Xaa Gln Asp Leu Lys

```
55
Leu Thr Ser Glu Glu Glu Ser Gln Arg Leu Lys Gly Ser Glu Asn Ser
                         70
Gln Pro Glu Glu Met Ser Gln Glu Pro Glu Ile Asn Xaa Gly Gly Asp
                     85
                                      90
Arg Lys Val Glu Xaa Xaa Met Lys Lys His Gly Ser Xaa His Met Gly
                 100
                                105
Phe Pro Xaa Asn Leu Xaa Asn Gly Ala Thr Ala Asp Asn Gly Asp Asp
                                120
              115
Gly Leu Ile Pro Pro Xaa Lys Xaa Xaa Thr Pro Glu Ser Xaa Gln Phe
                         135
Pro Asp Thr Glu Asn Glu Gln Tyr His Arg Asp Phe Ser Gly His Pro
     145 150
Xaa Phe Pro Thr Thr Leu Pro Ile Lys Gln
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<210> 492
<211> 216
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -15..-1

<400> 492 Met Val Cys Val Leu Val Leu Ala Ala Ala Ala Gly Ala Val Ala Val

-10 -5 Phe Leu Ile Leu Arg Ile Trp Val Val Leu Arg Ser Met Asp Val Thr Pro Arg Glu Ser Leu Ser Ile Leu Val Val Ala Gly Ser Gly Gly His 25 Thr Thr Glu Ile Leu Arg Leu Leu Gly Ser Leu Ser Asn Ala Tyr Ser 40 Pro Arg His Tyr Val Ile Ala Asp Thr Asp Glu Met Ser Ala Asn Lys 60 Ile Asn Ser Phe Glu Leu Xaa Arg Xaa Asp Arg Xaa Pro Ser Asn Met 75 Xaa Thr Lys Tyr Tyr Ile His Arg Ile Pro Xaa Ser Arg Glu Val Gln 90 Gln Ser Trp Pro Ser Thr Val Xaa Thr Thr Leu His Ser Met Trp Leu 105 Ser Xaa Pro Leu Ile His Arg Val Lys Pro Xaa Leu Val Leu Cys Asn 120 Gly Pro Gly Thr Cys Val Pro Ile Cys Val Ser Ala Leu Leu Leu Gly 135 140 Ile Leu Gly Ile Lys Lys Val Ile Ile Val Tyr Val Glu Ser Ile Cys 155 150 Arg Val Lys Thr Leu Ser Met Ser Gly Lys Ile Leu Phe His Leu Ser 165 170 Asn Tyr Phe Ile Val Gln Trp Pro Ala Leu Lys Glu Lys Tyr Pro Lys 180 185 Ser Val Tyr Leu Gly Arg Ile Val 200 195

<210> 493 <211> 134 <212> PRT

<221> SIGNAL <222> -29..-1

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<213> Homo sapiens
<220>
<221> SIGNAL
<222> -19..-1
<400> 493
Met Pro Leu Gly Ala Arg Ile Leu Phe His Gly Val Phe Tyr Ala Gly
                -15
                                    -10
Gly Phe Ala Ile Val Tyr Tyr Leu Ile Gln Lys Phe His Ser Arg Thr
Leu Tyr Tyr Lys Leu Ala Val Glu Gln Leu Gln Xaa His Pro Glu Ala
                        20
Gln Glu Ala Leu Gly Pro Pro Leu Asn Ile His Tyr Leu Lys Leu Ile
                    35
Asp Arg Glu Asn Phe Val Asp Ile Val Xaa Ala Lys Leu Lys Ile Pro
                                    55
Val Ser Gly Ser Lys Ser Glu Gly Leu Leu Tyr Val His Ser Ser Arg
                                70
Gly Gly Pro Phe Gln Arg Trp His Leu Asp Glu Val Phe Leu Glu Leu
Lys Asp Gly Gln Gln Ile Pro Val Phe Lys Leu Ser Gly Glu Asn Gly
Asp Glu Val Lys Lys Glu
110
<210> 494
<211> 85
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -16..-1
<400> 494
Met Ala Val Thr Ala Leu Ala Ala Xaa Thr Trp Leu Gly Val Trp Gly
                        -10
Val Arg Thr Met Gln Ala Arg Gly Phe Gly Ser Asp Gln Ser Glu Asn
                                    10
Val Asp Arg Gly Ala Gly Ser Ile Arg Glu Ala Gly Gly Ala Phe Gly
                                25
Lys Arg Glu Gln Ala Glu Glu Glu Arg Tyr Phe Arg Ala Gln Ser Thr
                            40
Glu Gln Leu Ala Xaa Leu Lys Lys Xaa His Glu Glu Glu Ile Val His
                                            60
His Arg Glu Gly Asp
<210> 495
<211> 292
<212> PRT
<213> Homo sapiens
<220>
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<400> 495
Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn His Thr Phe
                                   -20
               -25
Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr Glu Tyr Thr
                               -5
Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser Leu His Tyr
                                          15
Leu Leu Leu Pro Tyr Leu Leu Leu Gly Val Asn Leu Phe Phe Phe Thr
                   25
Leu Thr Cys Gly Thr Asn Pro Gly Ile Ile Thr Lys Ala Asn Glu Leu
                                   45
               40
Leu Phe Leu His Val Tyr Glu Phe Asp Glu Xaa Met Phe Pro Lys Asn
                               60
Val Arg Cys Ser Thr Cys Asp Leu Arg Lys Pro Ala Arg Ser Xaa His
                           75
Cys Xaa Val Cys Asn Trp Cys Val His Arg Phe Xaa His His Cys Val
                      90
Trp Val Asn Asn Cys Ile Gly Ala Trp Asn Ile Arg Xaa Phe Leu Ile
                                       110
                  105
Tyr Val Leu Thr Leu Thr Ala Ser Ala Ala Thr Val Ala Ile Val Ser
                                   125
               120
Thr Thr Phe Leu Val His Leu Val Val Met Ser Asp Leu Tyr Gln Glu
                               140
           135
Thr Tyr Ile Asp Asp Leu Gly His Leu His Val Met Asp Thr Val Phe
                          155
Leu Ile Gln Tyr Leu Phe Leu Thr Phe Pro Arg Ile Val Phe Met Leu
                                           175
                        170
Gly Phe Val Val Val Leu Xaa Phe Leu Leu Gly Gly Tyr Leu Leu Phe
                                       190
                   185
Val Leu Tyr Leu Ala Ala Thr Asn Gln Thr Thr Asn Glu Trp Tyr Arg
               200
                                   205
Xaa Asp Trp Ala Trp Cys Gln Arg Cys Pro Leu Val Ala Trp Pro Pro
                               220
Ser Ala Glu Pro Gln Val His Arg Asn Ile His Ser His Gly Leu Arg
                           235
Xaa Asn Leu Gln Glu Ile Phe Leu Pro Ala Phe Pro Cys His Glu Arg
                       250
Lys Lys Gln Glu
260
```

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<210> 496
<211> 122
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -56..-1
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 Phe Cys Ala Ser Phe Pro Ser Gly Xaa Leu Ser Pro Pro Gly Pro Leu

 25
 30
 35
 40

 Pro Gly Val Arg Gly Leu Pro Leu Pro Ser Val Phe Tyr Ser Cys Gly
 50
 55

 Ala His Pro Lys Val Leu Lys Val Ala Leu
 65
 65

<211> 99 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -13..-1 <400> 498 Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg Arg Pro -10 -5 Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu Ala His 10 Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys Trp Arg Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn Ser Ser 40 45 Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr Pro Met Arg Arg Ser Ser Cys His Leu Glu Cys Xaa Val Ile Phe Leu Leu Gly Arg Gln Leu 85

<210> 499 <211> 99 <212> PRT <213> Homo sapiens

<210> 498

<220>

<221> SIGNAL

<222> -13..-1

<400> 499

Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg Arg Pro
-10 -5 1

Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu Ala His
5 10 15

Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys Trp Arg 20 25 30 35

Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn Ser Ser 40 45 50

Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr Pro Met 55 60 65

Arg Arg Ser Ser Cys His Leu Xaa Cys Gln Val Ile Phe Leu Leu Gly
70 75 80

Arg Gln Leu 85

<210> 500

<211> 108

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -25..-1

<400> 500

Met Ser Leu Thr Ser Ser Ser Ser Val Arg Val Glu Trp Ile Ala Ala
-25 -20 -15 -10

Val Thr Ile Ala Ala Gly Thr Ala Ala Ile Gly Tyr Leu Ala Tyr Lys
-5 1 5

Arg Phe Tyr Val Lys Asp His Arg Asn Lys Ala Met Ile Asn Leu His
10 15 20

Ile Gln Lys Asp Asn Pro Lys Ile Val His Ala Phe Asp Met Glu Asp
25 30 35

Leu Gly Asp Lys Ala Val Tyr Cys Arg Cys Trp Arg Ser Lys Lys Phe 40 45 50 50

Pro Phe Cys Asp Gly Ala His Thr Lys His Asn Glu Glu Thr Gly Asp
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Asn Val Gly Pro Leu Ile Ile Lys Lys Glu Thr
75 80

<210> 501

<211> 183

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -15..-1

<400> 501

Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu Ala Val Leu Ala Trp
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Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg Met Lys Ser Arg Glu

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10
Gln Gly Arg Arg Leu Gly Ala Glu Ser Arg Thr Leu Leu Val Ile Ala
                        25
His Pro Asp Asp Glu Ala Met Phe Phe Ala Pro Thr Val Leu Gly Leu
Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys Phe Ser Ala Gly Asn
                                       60
Tyr Tyr Asn Gln Gly Glu Thr Arg Lys Lys Glu Leu Leu Gln Ser Cys
Asp Val Leu Gly Ile Pro Leu Ser Ser Val Met Ile Ile Asp Asn Arg
                               90
Asp Phe Pro Xaa Asp Pro Gly Met Gln Trp Asp Thr Xaa His Val Ala
                           105
                                               110
Xaa Val Leu Leu Gln His Ile Glu Val Asn Gly Ile Asn Leu Val Val
                       120
                                           125
Thr Phe Asp Ala Gly Gly Xaa Ser Gly His Ser Asn His Ile Ala Leu
                   135
                                      140
Tyr Ala Ala Val Arg Lys Leu Glu Gly Gln Ile Cys Lys Pro Cys Gly
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                                   155
Thr Gly Gln Asp Phe Lys Glu
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 Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu Ala Val Leu Ala Trp -15
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 -5
 1

 Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg Met Lys Ser Arg Glu 5
 10
 15

 Gln Gly Xaa Arg Leu Gly Ala Glu Ser Arg Thr Leu Leu Val Ile Ala 20
 25
 30

 His Pro Asp Asp Asp Glu Ala Met Phe Phe Ala Pro Thr Val Leu Gly Leu 35
 40
 45

 Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys Phe Ser Ala Val Phe 50
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 60
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 Arg Arg Glu Leu Ser Glu Tyr Thr Glu Xaa Leu Thr Ser Glu Pro Leu 70
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<210> 503 <211> 183 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -57..-1

Xaa Ala

<400> 503
Met Asp Val Thr Gly Asp Glu Glu Glu Glu Ile Lys Gln Glu Ile Asn
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-50
-45

Met Leu Lys Lys Tyr Ser His His Arg Asn Ile Ala Thr Tyr Tyr Gly -35 Ala Phe Ile Lys Lys Asn Pro Pro Gly Met Asp Asp Gln Leu Trp Leu -20 -15 Val Met Glu Phe Cys Gly Ala Gly Ser Val Thr Asp Leu Ile Lys Asn Thr Lys Gly Asn Thr Leu Lys Glu Glu Trp Ile Ala Tyr Ile Cys Xaa 15 Glu Ile Leu Arg Gly Leu Xaa His Leu His Gln His Lys Val Ile His Arg Xaa Ile Lys Gly Gln Asn Val Leu Leu Thr Glu Asn Ala Glu Val Lys Leu Val Asp Phe Gly Xaa Xaa Ala Gln Leu Asp Arg Thr Val Gly 65 Arg Xaa Asn Thr Phe Ile Gly Thr Pro Tyr Trp Met Ala Pro Xaa Val 80 Ile Ala Cys Asp Glu Asn Pro Xaa Ala Thr Tyr Asp Phe Lys Xaa Asp 95 Leu Trp Ser Leu Gly Ile Thr Ala Ile Glu Met Ala Glu Gly Leu Pro 110 Leu Ser Val Thr Cys Thr Pro

<210> 504 <211> 140 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

<400> 504

<222> -14..-1

Met Phe Leu Thr Ala Leu Leu Trp Arg Gly Arg Ile Pro Gly Arg Gln ~10 Trp Ile Gly Lys His Arg Arg Pro Arg Phe Val Ser Leu Arg Ala Lys 10 Gln Asn Met Ile Arg Arg Leu Glu Ile Glu Ala Glu Asn His Tyr Trp 25 Leu Ser Met Pro Tyr Met Thr Arg Glu Gln Glu Arg Gly His Ala Ala Leu Arg Arg Arg Glu Ala Phe Glu Ala Ile Lys Ala Ala Ala Thr Ser 60 Lys Phe Pro Pro His Arg Phe Ile Ala Asp Gln Leu Asp His Leu Asn 75 Xaa His Gln Glu Met Val Leu Ile Leu Ser Arg His Pro Trp Ile Leu 90 Trp Ile Thr Glu Leu Thr Ile Phe Thr Trp Ser Gly Leu Lys Asn Cys 105 Ser Leu Cys Glu Asn Glu Leu Trp Thr Ser Leu Tyr 120

<210> 505 <211> 59 <212> PRT <213> Homo sapiens

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                                  -5
Cys Ser Ala Xaa Leu Gly Arg Ala Ala Ser Gly Xaa Tyr Ser Arg Asn
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Trp Leu Pro Thr Pro Pro Ala Thr Gly Pro Leu Pro Ser Ser Gln Thr
                   25
Gly His Met Arg Met Ala Ala Leu Leu Pro Gln
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<210> 506
<211> 101
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -36..-1
<400> 506
Met Gly Pro Tyr Asn Val Ala Val Pro Ser Asp Val Ser His Ala Arg
                       -30
                                          -25
Phe Tyr Phe Leu Phe His Arg Pro Leu Arg Leu Leu Asn Leu Leu Ile
                   -15
                                      -10 .
Leu Ile Glu Gly Ser Val Val Phe Tyr Gln Leu Tyr Ser Leu Leu Arg
                    5
               1
                                         10
Ser Glu Lys Trp Asn His Thr Leu Ser Met Ala Leu Ile Leu Phe Cys
    15
                           20
Asn Tyr Tyr Val Leu Phe Lys Leu Leu Arg Asp Arg Xaa Xaa Leu Gly
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Arg Ala Tyr Ser Tyr Pro Leu Asn Ser Tyr Glu Leu Lys Ala Asn Xaa
Ala Ala Ser Xaa Gln
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<211> 341
<212> PRT
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                                       -45
Ala Leu Cys Lys Arg Leu Leu Ala Glu Asp Asp Glu Leu His Leu Cys
               -35
                                   -30
Leu Ala Cys Arg Asn Met Ser Lys Ala Glu Ala Val Cys Ala Ala Leu
           -20
                              -15
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Leu Ala Ser His Pro Thr Ala Glu Val Thr Ile Val Gln Val Asp Val

Ser Asm Leu Gln Ser Phe Phe Arg Ala Ser Lys Glu Leu Lys Gln Arg

20

15

WO 99/31236 -354 - PCT/IB98/02122

Phe Gln Arg Leu Asp Cys Ile Tyr Leu Asn Ala Gly Ile Met Pro Asn 35 Pro Gln Leu Asn Ile Lys Ala Leu Phe Phe Gly Leu Phe Ser Arg Lys 50 Val Ile His Met Phe Ser Thr Ala Glu Gly Leu Leu Thr Gln Gly Asp Lys Ile Thr Ala Asp Gly Leu Gln Glu Val Phe Glu Thr Asn Val Phe Gly His Phe Ile Leu Ile Arg Glu Leu Glu Pro Leu Leu Cys His Ser 100 Asp Asn Pro Ser Gln Leu Ile Trp Thr Ser Ser Arg Ser Ala Arg Lys 110 115 Ser Asn Phe Ser Leu Glu Asp Phe Gln His Ser Lys Gly Lys Glu Pro 130 Tyr Ser Ser Ser Lys Tyr Ala Thr Asp Leu Leu Ser Val Ala Leu Asn 145 Arg Asn Phe Asn Gln Gln Gly Leu Tyr Ser Asn Val Ala Cys Pro Gly 160 Thr Ala Leu Thr Asn Leu Thr Tyr Gly Ile Leu Pro Pro Phe Ile Trp 175 180 Thr Leu Leu Met Pro Ala Ile Leu Leu Leu Arg Phe Phe Ala Asn Ala 195 190 Phe Thr Leu Thr Pro Tyr Asn Gly Thr Glu Ala Leu Val Trp Leu Phe 210 His Gln Lys Pro Glu Ser Leu Asn Pro Leu Ile Lys Tyr Leu Ser Ala 225 Thr Thr Gly Phe Gly Arg Asn Tyr Ile Met Thr Gln Lys Met Asp Leu 240 Asp Glu Asp Thr Ala Glu Lys Phe Tyr Gln Lys Leu Leu Glu Leu Glu 255 260 Lys His Ile Arg Val Thr Ile Gln Lys Thr Asp Asn Gln Ala Arg Leu 270 275 Ser Gly Ser Cys Leu 285

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<222> -26..-1
<400> 509
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Thr Asn Gln Val Leu Ile Thr Ala Arg Ala Val Pro Thr Lys Lys Ala
                                        3
Ser Val Arg Cys Val Glu Lys Arg Phe Trp Ile Pro Lys Thr Thr Ser
            10
Lys His Leu Ser Arg Cys Ile Asp Gly Ile Ser Gly Phe Leu Asn Asp
                            30
Phe Thr Phe Cys Leu Glu Phe Ser Arg His Arg Cys Gln Leu Thr Glu
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<211> 158
<212> PRT
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<222> -44..-1
<400> 510
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                -40
                                    -35
Asp Trp Ser Glu Arg Arg Asn Ala Val Ala Ser Val Val Ala Gly Ile
                                -20
Leu Phe Phe Thr Gly Trp Trp Ile Met Ile Asp Ala Ala Val Val Tyr
                            -5
Pro Lys Pro Glu Gln Leu Asn His Ala Phe His Thr Cys Gly Val Phe
                    10
                                        15
Ser Thr Leu Ala Phe Phe Met Ile Asn Ala Val Ser Asn Ala Gln Val
                                    30
Arg Gly Asp Ser Tyr Glu Ser Gly Cys Leu Gly Arg Thr Gly Ala Arg
                                45
Val Trp Leu Phe Ile Gly Phe Met Leu Met Phe Gly Ser Leu Ile Ala
                            60
Ser Met Trp Ile Leu Phe Gly Ala Tyr Val Thr Gln Asn Thr Asp Val
Tyr Pro Gly Leu Ala Val Phe Phe Gln Asn Ala Leu Ile Phe Phe Ser
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Thr Leu Ile Tyr Lys Phe Gly Arg Thr Glu Glu Leu Trp Thr
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<212> PRT

<213> Homo sapiens

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<400> 511
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Asn Gly Asn Leu Lys Glu Lys Asp Ile Leu Val Leu Pro Leu Asp Leu
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Thr Asp Thr Gly Ser His Glu Ser Gly Tyr Gln Ser Cys Ser Pro Gly 85 90 95 100 Ile Trp

iic iip

<210> 512 <211> 199 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -62..-1

<400> 512

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-60
-55
-50
Yee Leu The Cys Cys

Xaa Leu Ile Glu Leu Asn Tyr Leu Gly Thr Val Ser Leu Thr Lys Cys
-45 -40 -35

Val Leu Pro His Met Ile Glu Arg Lys Gln Gly Lys Ile Val Thr Val
-30 -25 -20 -15

Asn Ser Ile Leu Gly Ile Ile Ser Val Pro Leu Ser Ile Gly Tyr Cys
-10 -5 1

Ala Ser Lys His Ala Leu Arg Gly Phe Phe Asn Gly Leu Arg Thr Glu 5 10 15

Leu Ala Thr Tyr Pro Gly Ile Ile Val Ser Asn Ile Cys Pro Gly Pro 20 25 30

Val Gln Ser Asn Ile Val Glu Asn Ser Leu Ala Gly Glu Val Thr Lys 35 40 45 50

Thr Ile Gly Asn Asn Gly Asn Gln Ser His Lys Met Thr Thr Ser Arg
55 60 65

Cys Val Arg Leu Met Leu Ile Ser Met Ala Asn Asp Leu Lys Glu Val

Trp Ile Ser Glu Gln Pro Phe Leu Leu Val Thr Tyr Leu Trp Gln Tyr 85 90 95

Met Pro Thr Trp Ala Trp Trp Ile Thr Asn Lys Met Gly Lys Lys Arg 100 105 110

Ile Glu Asn Phe Lys Ser Gly Val Asp Ala Xaa Ser Ser Tyr Phe Lys
115 120 125 130

Ile Phe Lys Thr Lys His Asp

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<211> 180
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
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<400> 513
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                                       -15
Ser Val Met Ala Ala Leu Thr Phe Gly Cys Phe Ile Xaa Thr Ala Phe
        -5
Lys Asp Arg Ser Val Pro Val Arg Leu His Val Ser Arg Ile Met Leu
                           15
Lys Asn Val Glu Asp Phe Thr Gly Pro Arg Glu Arg Ser Asp Leu Gly
Phe Ile Thr Phe Asp Ile Thr Ala Asp Leu Glu Asn Ile Phe Asp Trp
                   45
Asn Val Lys Gln Leu Phe Leu Tyr Leu Ser Ala Glu Tyr Ser Thr Lys
                                   65
Asn Asn Ala Leu Asn Gln Xaa Val Leu Trp Asp Lys Ile Val Leu Arg
                               80
Gly Asp Asn Pro Lys Leu Leu Leu Lys Asp Met Lys Thr Lys Tyr Phe
                           95
Phe Phe Asp Asp Gly Asn Gly Leu Xaa Gly Asn Arg Asn Val Thr Leu
                       110
Thr Leu Ser Trp Asn Val Val Pro Asn Ala Gly Ile Leu Pro Leu Val
                   125
                                      130
Thr Gly Ser Gly His Val Ser Val Pro Phe Pro Asp Thr Tyr Glu Ile
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Thr Lys Ser Tyr
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<210> 514 <211> 120 <212> PRT <213> Bos taurus

115

<400> 514

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 Thr
 Gly
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 Gln
 Gly
 Arg
 Ala
 Thr
 Phe
 Gln
 Phe
 Leu
 Pro
 Asp

 Glu
 Ala
 Arg
 Ser
 Leu
 Pro
 Pro
 Pro
 Leu
 Thr
 Asp
 Pro
 Arg
 Leu
 Ala
 Ala
 Asp
 Pro
 Arg
 Arg

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                                                                  180
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ggaggaggcg cggcaagacg tggaggccct cctgagccgc acggtcagaa ctcagatact
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qaccggcaag gagctccgag ttgccaccca ggaaaaaagag ggctcctctg ggagatgtat
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                                                                  360
gettactete ttaggeettt catteatett ggeaggaett attgttggtg gageetgeat
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                                                                  600
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                                                                  780
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tattqttqaq accaaqatct gtcaagagta agaggcaaca gatagagtgt ccttggtaat
                                                                  960
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1082
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                                                                  180
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                                                                  300
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atagaaatcc ttaaaaaagt aagccaacaa acagaaaaga caacaacagt gaaattcaac
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agaagaaaag taatggactc tgatgaagat gacgattatt gaactacaag tgctcacaga
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His Val Gly Asp Ser Ala Leu Met Gly Cys Val Phe Gln Ser Thr Glu
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35

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<400> 518

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